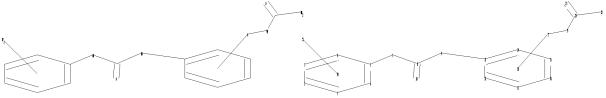
```
5-7 7-8 8-9 8-10 9-13 16-17 17-18 18-19 19-20 19-21 21-26
exact bonds :
   1-29 2-27 3-22 4-28 6-30 11-34 12-33 14-31 15-32
normalized bonds:
   1-2 1-6 2-3 3-4 4-5 5-6 11-12 11-16 12-13 13-14 14-15 15-16
isolated ring systems :
   containing 1 : 11 :
G1:H,CH3
Match level :
   1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:Atom
   12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:CLASS 18:Atom 19:CLASS 20:CLASS 21:CLASS
   22:CLASS 26:CLASS 27:CLASS 28:CLASS 29:CLASS 30:CLASS 31:CLASS 32:CLASS 33:CLASS
   34:CLASS
Generic attributes :
   18:
   Saturation
                         : Unsaturated
   Number of Carbon Atoms : less than 7
   Number of Hetero Atoms : Exactly 1
   Type of Ring System
                      : Monocyclic
Element Count :
   Node 18: Limited
```

C,C5 N,N1 O,O0 S,S0 =>

Uploading C:\Program Files\Stnexp\Queries\09993647.str



```
chain nodes :
7 8 9 10 17 19 20 21 22 23
ring nodes :
1 2 3 4 5 6 11 12 13 14 15 16
chain bonds :
5-7 7-8 8-9 8-10 9-13 17-19 19-20 20-21 20-22
ring bonds :
    1-6 2-3
              3-4 4-5 5-6 11-12 11-16 12-13 13-14 14-15 15-16
exact/norm bonds :
5-7 7-8 8-9 8-10 9-13 17-19 19-20 20-21 20-22
normalized bonds :
1-2 \quad 1-6 \quad 2-3 \quad 3-4 \quad 4-5 \quad 5-6 \quad 11-12 \quad 11-16 \quad 12-13 \quad 13-14 \quad 14-15 \quad 15-16
isolated ring systems :
containing 1 : 11 :
Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS
 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:CLASS 18:Atom 19:Atom
20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:Atom
Generic attributes :
19:
Saturation
                       : Unsaturated
Number of Carbon Atoms : less than 7
Number of Hetero Atoms : Exactly 1
Type of Ring System
                       : Monocyclic
Element Count :
Node 19: Limited
    C,C5
   N,N1
    0,00
    S,S0
```

### L1 STRUCTURE UPLOADED

=>

Uploading C:\Program Files\Stnexp\Queries\09993647 (a).str



```
chain nodes :
7  8  9  10  17  19  20  21  22  23  28
ring nodes :
1  2  3  4  5  6  11  12  13  14  15  16
chain bonds :
5-7  7-8  8-9  8-10  9-13  17-19  19-20  20-21  20-22  22-28
ring bonds :
1-2  1-6  2-3  3-4  4-5  5-6  11-12  11-16  12-13  13-14  14-15  15-16
exact/norm bonds :
5-7  7-8  8-9  8-10  9-13  17-19  19-20  20-21  20-22  22-28
normalized bonds :
1-2  1-6  2-3  3-4  4-5  5-6  11-12  11-16  12-13  13-14  14-15  15-16
isolated ring systems :
containing 1 : 11 :
```

# G1:H,CH3

Match level: 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:CLASS 18:Atom 19:Atom 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:Atom 28:CLASS Generic attributes : 19: Saturation : Unsaturated Number of Carbon Atoms : less than 7 Number of Hetero Atoms : Exactly 1 Type of Ring System : Monocyclic Element Count : Node 19: Limited C,C5 N,N1 0,00 S,S0 L2 STRUCTURE UPLOADED

=> d 12

L2 HAS NO ANSWERS

STR

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

Structure attributes must be viewed using STN Express query preparation.

=> s 12 sss sam

SAMPLE SEARCH INITIATED 07:48:33 FILE 'REGISTRY'

839 TO ITERATE SAMPLE SCREEN SEARCH COMPLETED -

100.0% PROCESSED 839 ITERATIONS 43 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 15043 TO 18517 PROJECTED ANSWERS: 467 TO 1253

43 SEA SSS SAM L2 L3

=> => ....Testing the current file.... screen

ENTER SCREEN EXPRESSION OR (END):end

=> screen 1840

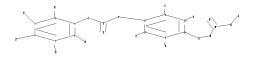
L4 SCREEN CREATED

=> screen 2016 OR 2026 OR 1841 OR 2039 OR 2040 OR 2045 OR 2047

L5 SCREEN CREATED

=>

Uploading C:\Program Files\Stnexp\Queries\09993647 (b).str



chain nodes :
7 8 9 10 17 18 19 20 21 22 26 27 28 29 30 31 32 33 34
ring nodes :
1 2 3 4 5 6 11 12 13 14 15 16
chain bonds :
1-29 2-27 3-22 4-28 5-7 6-30 7-8 8-9 8-10 9-13 11-34 12-33 14-31 15-32
16-17 17-18 18-19 19-20 19-21 21-26
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 11-12 11-16 12-13 13-14 14-15 15-16
exact/norm bonds :
5-7 7-8 8-9 8-10 9-13 16-17 17-18 18-19 19-20 19-21 21-26
exact bonds :

```
1-29 \quad 2-27 \quad 3-22 \quad 4-28 \quad 6-30 \quad 11-34 \quad 12-33 \quad 14-31 \quad 15-32
normalized bonds :
1-2 \quad 1-6 \quad 2-3 \quad 3-4 \quad 4-5 \quad 5-6 \quad 11-12 \quad 11-16 \quad 12-13 \quad 13-14 \quad 14-15 \quad 15-16
isolated ring systems :
containing 1 : 11 :
G1:H, CH3
Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:CLASS 18:Atom 19:CLASS
20:CLASS 21:CLASS 22:CLASS 26:CLASS 27:CLASS 28:CLASS 29:CLASS 30:CLASS
31:CLASS 32:CLASS 33:CLASS 34:CLASS
Generic attributes :
18:
Saturation
                        : Unsaturated
Number of Carbon Atoms : less than 7
Number of Hetero Atoms : Exactly 1
Type of Ring System : Monocyclic
Element Count :
Node 18: Limited
    C,C5
    N,N1
    0,00
    S,SO
L6
       STRUCTURE UPLOADED
=> que L6 AND L4 NOT L5
L7 QUE L6 AND L4 NOT L5
=> d 17
L7 HAS NO ANSWERS
                 SCR 1840
T<sub>1</sub>5
                 SCR 2016 OR 2026 OR 1841 OR 2039 OR 2040 OR 2045 OR 2047
                 STR
L6
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
Structure attributes must be viewed using STN Express query preparation.
                 QUE L6 AND L4 NOT L5
L7
\Rightarrow s 17 sss sam
SAMPLE SEARCH INITIATED 07:53:29 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 29 TO ITERATE
100.0% PROCESSED
                   29 ITERATIONS
                                                                      4 ANSWERS
SEARCH TIME: 00.00.01
```

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 257 TO 903 PROJECTED ANSWERS: 4 TO 200

L8 4 SEA SSS SAM L6 AND L4 NOT L5

=> => s 17 sss ful

FULL SEARCH INITIATED 07:54:19 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 509 TO ITERATE

100.0% PROCESSED 509 ITERATIONS 78 ANSWERS

SEARCH TIME: 00.00.01

L9 78 SEA SSS FUL L6 AND L4 NOT L5

=> => s 19

L10 1515 L9

=> s bayer?

L11 10038 BAYER?

=> s 110 and 111

L12 11 L10 AND L11

=> d 112 1-11 bib, ab, hitstr

L12 ANSWER 1 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN

AN 2010:317457 CAPLUS

DN 153:471241

TI Palliative management of hepatocarcinoma with sorafenib (Nexavar). Results of the SHARP study (Sorafenib Hepatocarcinoma Assessment Randomized Protocol trial)

AU Detry, O.; Delwaide, J.; De Roover, A.; Meunier, P.; Van Daele, D.; Lamproye, A.; Honore, P.; Polus, M.

CS Service de Chirurgie Abdominal et Transplantation, CHU de Liege, Belg.

SO Revue Medicale de Liege (2009), 64(3), 168-170 CODEN: RMLIAC; ISSN: 03 0-629X

PB Revue Medicale de Liege

DT Journal; General Review

LA French

AΒ A review. Curative management of early-stage hepatocarcinoma may include partial hepatic resection, liver transplantation or tumoral necrosis using radiofrequency ablation or alcoholisation. Until recently, no efficient therapeutic mean was available for advanced hepatocarcinoma. Sorafenib (Nexavar, Bayer) is a multikinase inhibitor that decreases tumoral proliferation and angiogenesis, and increases apoptosis in many cancer models. The results of a phase 3 randomized, multicentric, study, entitled SHARP, have now demonstrated that sorafenib increases survival in patients with advanced hepatocarcinoma developed in Child A cirrhosis. Mean survival gain was a little less than 3 mo, without any radiol. response or improvement in the delay before symptomatic progression of the disease. The monthly cost of sorafenib is a little more than 5,000 euros. It is now crucial to evaluate the potential role of sorafenib in adjuvant therapy after liver resection or radiofrequency ablation of hepatocarcinoma. The CHU of Liege is taking part to a randomized, multicentric study evaluating the use of sorafenib after liver resection or radiofrequency ablation for hepatocarcinoma. Another future evaluation could be the association of sorafenib with other antitumoral agents.

IT 284461-73-0, Sorafenib 475207-59-1, Nexavar RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (palliative management of hepatocarcinoma with sorafenib)

RN 284461-73-0 CAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 475207-59-1 CAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 284461-73-0

CMF C21 H16 C1 F3 N4 O3

CM 2

CRN 104-15-4 CMF C7 H8 O3 S

L12 ANSWER 2 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN

AN 2009:1629506 CAPLUS

DN 153:162840

TI Platelet count less than SHARP: what does a case series reveal?

AU Saif, M. Wasif

CS Section of Medical Oncology, Yale University School of Medicine, New Haven, CT, 06520, USA

SO Expert Opinion on Drug Safety (2010), 9(1), 1-8 CODEN: EODSA9; ISSN: 1474-0338

PB Informa Healthcare

DT Journal; General Review

LA English

AΒ Hepatocellular carcinoma (HCC) is increasing in nos. worldwide, A review. and no effective systemic treatment existed for advanced HCC until SHARP (Sorafenib in HCC Assessment Randomized Protocol) study proved sorafenib (Nexavar, Bayer Pharmaceuticals, Wayne, NJ, USA) prolonged survival vs. placebo. Child-Pugh class A liver function and a platelet count of  $\geq$  60,000/mm3 were among the inclusion criteria for SHARP. No safety data in patients with < 60,000/mm3 of platelets are present. Thrombocytopenia is one of the most frequent challenges faced in patients with chronic liver diseases. We report a series of three patients with HCC and platelet count < 60,000/mm3 who were successfully treated with sorafenib with no complications. We describe the current data on sorafenib and challenges faced in patients with HCC. In addition, we emphasize the need for informed consent when facing factors that predispose to bleeding (esophageal varices, coagulopathy and thrombocytopenia), possible band ligation before the start of sorafenib, careful clin. monitoring and discontinuation of sorafenib when major bleeding occurs.

IT 475207-59-1, Nexavar

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (Nexavar was safe and effective but reduced platelet count in patient with hepatocellular carcinoma)

RN 475207-59-1 CAPLUS

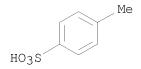
CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 284461-73-0 CMF C21 H16 C1 F3 N4 O3

CM 2

CRN 104-15-4 CMF C7 H8 O3 S



RE.CNT 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
L12 ANSWER 3 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN
```

AN 2009:324660 CAPLUS

DN 151:235838

TI Sorafenib for the treatment of unresectable hepatocellular carcinoma

- AU Kane, Robert C.; Farrell, Ann T.; Madabushi, Rajanikanth; Booth, Brian; Chattopadhyay, Somesh; Sridhara, Rajeshwari; Justice, Robert; Pazdur, Richard
- CS Office of Oncology Drug Products, Center for Drug Evaluation and Research, U.S. Food and Drug Administration, Silver Spring, MD, USA
- SO Oncologis (2009) 14(1), 95-100 CODEN: OCOLF6; ISSN: 1083-7159
- PB AlphaMed Press
- DT Journal
- LA English
- Purpose. To describe the U.S. Food and Drug Administration (FDA) review AΒ and approval of sorafenib (Nexavar; Bayer Pharmaceuticals Corp., Montville, NJ, and Onyx Pharmaceuticals Corp., Emeryville, CA), an oral kinase inhibitor, for the treatment of patients with unresectable hepatocellular carcinoma (HCC). Exptl. Design. The FDA independently analyzed an international, double-blind, placebo-controlled trial comparing the effect of best supportive care plus sorafenib or matching placebo on overall survival. Eligible patients had unresectable, biopsy-proven HCC and had not received prior systemic therapy. Results. Among the 602 randomized patients (placebo, 303; sorafenib, 299), baseline characteristics were well balanced, and 97% were Child-Pugh score A. HCC was "advanced" in 70% overall, as defined by extrahepatic metastases or by tumor radiog. visible in venous structures outside the liver. Underlying liver diseases included hepatitis B (18%), hepatitis C (28%), and alc.-related (26%). The trial was stopped following a prespecified second interim anal. showing a statistically significant survival advantage for sorafenib [median, 10.7 vs 7.9 mo; hazard ratio, 0.69 (95% confidence interval, (0.55, 0.87)), p = 0.00058]. Adverse events in sorafenib-treated patients included diarrhea in 55% (grade 3, 10%), hand-foot syndrome in 21% (grade 3,8%), rash in 19% (grade 3,1%), and cardiac ischemia or infarction in 2.7% (vs. 1.3% for placebo). On sorafenib, treatment-emergent hypertension occurred in 9% of patients (placebo, 4%) and was grade 3 in 4% (placebo, 1%); elevated serum lipase occurred in 40% (placebo, 37%); hypophosphatemia occurred in 35% (placebo, 11%). Conclusions. Sorafenib is the first systemic therapy to demonstrate a survival benefit in a randomized trial for unresectable HCC and has received FDA approval for this indication.
- IT 475207-59-1, Nexavar
  - RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (sorafenib for treating unresectable hepatocellular carcinoma)
- RN 475207-59-1 CAPLUS
- CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 284461-73-0

CMF C21 H16 C1 F3 N4 O3

CM 2

CRN 104-15-4 CMF C7 H8 O3 S

OSC.G 11 THERE ARE 11 CAPLUS RECORDS THAT CITE THIS RECORD (11 CITINGS)
RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L12 ANSWER 4 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN
- AN 2008:1542770 CAPLUS
- DN 151:48669
- TI Efficacy and safety of sorafenib in patients in the Asia-Pacific region with advanced hepatocellular carcinoma: a phase III randomized, double-blind, placebo-controlled trial
- AU Cheng, Ann-Lii; Kang, Yoon-Koo; Chen, Zhendong; Tsao, Chao-Jung; Qin, Shukui; Kim, Jun Suk; Luo, Rongcheng; Feng, Jifeng; Ye, Shenglong; Yang, Tsai-Sheng; Xu, Jianming; Sun, Yan; Liang, Houjie; Liu, Jiwei; Wang, Jiejun; Tak, Won Young; Pan, Hongming; Burock, Karin; Zou, Jessie; Voliotis, Dimitries, Guan, Zhongzhen
- CS National Taiwam University Hospital, Taipei, Taiwan
- SO Lancet Oncology (2009), 10(1), 25-34 CODEN: LOANBN; ISSN: 1470-2045
- PB Elsevier Ltd.
- DT Journal
- LA English
- AΒ Most cases of hepatocellular carcinoma occur in the Asia-Pacific region, where chronic hepatitis B infection is an important etiol. factor. Assessing the efficacy and safety of new therapeutic options in an Asia-Pacific population is thus important. We did a multinational phase III, randomized, double-blind, placebo-controlled trial to assess the efficacy and safety of sorafenib in patients from the Asia-Pacific region with advanced (unresectable or metastatic) hepatocellular carcinoma. Between Sept 20, 2005, and Jan 31, 2007, patients with hepatocellular carcinoma who had not received previous systemic therapy and had Child-Pugh liver function class A, were randomly assigned to receive either oral sorafenib (400 mg) or placebo twice daily in 6-wk cycles, with efficacy measured at the end of each 6-wk period. Eligible patients were stratified by the presence or absence of macroscopic vascular invasion or extrahepatic spread (or both), Eastern Cooperative Oncol. Group performance status, and geog. region. Randomization was done centrally and in a 2:1 ratio by means of an interactive voice-response system. There was no predefined primary endpoint; overall survival, time to progression (TTP), time to symptomatic progression (TTSP), disease control rate (DCR), and safety were assessed. Efficacy analyses were done by intention to treat. This trial is registered with, number Two hundred and seventy-one 271 patients from 23 centers in China, South Korea, and Taiwan were enrolled in the study. Of these, 226 patients were randomly assigned to the exptl. group (n=150) or to the placebo group (n=76). Median overall survival was 6.5 mo (95% CI 5.56-7.56) in patients treated with sorafenib, compared with 4.2 mo (3.75-5.46) in those who received placebo (hazard ratio [HR] 0.68 [95% CI 0.50-0.93]; p=0.014). Median TTP was 2.8 mo (2.63-3.58) in the sorafenib group compared with 1.4 mo (1.35-1.55) in the placebo group (HR 0.57 [0.42-0.79]; p=0.0005). The most frequently reported grade 3/4 drug-related adverse events in the 149 assessable patients treated with sorafenib were hand-foot skin reaction (HFSR; 16 patients [10.7%]), diarrhea (nine patients [6.0%]), and fatigue (five patients [3.4%]). The most common adverse events resulting in dose redns. were HFSR (17 patients [11.4%]) and diarrhea (11 patients [7.4%]); these adverse events rarely led to discontinuation. Sorafenib is effective for the treatment of advanced hepatocellular carcinoma in patients from the Asia-Pacific region, and is well tolerated. Taken together with data from the Sorafenib Hepatocellular Carcinoma Assessment Randomized Protocol (SHARP) trial, sorafenib seems to be an appropriate option for the treatment of advanced hepatocellular carcinoma. Funding: Bayer HealthCare Pharmaceuticals and Onyx Pharmaceuticals, Inc.

IT 284461-73-0, Sorafenib

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(sorafenib was well tolerated and effective in treatment of patient with metastatic hepatocellular carcinoma in Asia-Pacific region)

RN 284461-73-0 CAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

OSC.G 102 THERE ARE 102 CAPLUS RECORDS THAT CITE THIS RECORD (102 CITINGS)

RE.CNT 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L12 ANSWER 5 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN
- AN 2008:798022 CAPLUS
- DN 149:263859
- TI Dissecting and Targeting the Growth Factor-Dependent and Growth Factor-Independent Extracellular Signal-Regulated Kinase Pathway in Human Schwannoma
- AU Ammoun, Sylwia; Flaiz, Christine; Ristic, Natalia; Schuldt, Jennifer; Hanemann, C. Oliver
- CS Clinical Neurobiology, Peninsula College for Medicine and Dentistry, Plymouth, PL6 8BU
- SO Cancer Research (2008), 68(13), 5236-5245 CODEN: CNREA8; 1980, 0008-5472
- PB American Association for Cancer Research
- DT Journal
- LA English
- AΒ Schwannomas are tumors of the nervous system that occur sporadically and in patients with the cancer predisposition syndrome neurofibromatosis type 2 (NF2). Schwannomas and all NF2-related tumors are caused by loss of the tumor suppressor merlin. Using our human in vitro model for schwannoma, we analyzed extracellular signal-regulated kinase 1/2 (ERK1/2) and AKT signaling pathways, their upstream growth factor receptors, and their role in schwannoma cell proliferation and adhesion to find new systemic therapies for these tumors that, to date, are very difficult to treat. show here that human primary schwannoma cells show an enhanced basal Raf/mitogen-activated protein/ERK kinase/ERK1/2 pathway activity compared with healthy Schwann cells. Due to a strong and prolonged activation of platelet-derived growth factor receptor  $\beta$  (PDGFR $\beta$ ), which is highly overexpressed, ERK1/2 and AKT activation was further increased in schwannoma, leading to increased proliferation. Using specific inhibitors, we discovered that ERK1/2 activation involves the integrin/focal adhesion kinase/Src/Ras signaling cascades and PDGFR $\beta$ -mediated ERK1/2 activation is triggered through the phosphatidylinositol 3-kinase/protein kinase C/Src/c-Raf pathway. the complexity of signals leading to schwannoma cell proliferation, potential new therapeutic agents should target several signaling pathways. The PDGFR and c-Raf inhibitor sorafenib (BAY 43-9006; Bayer Pharmaceuticals), currently approved for treatment of advanced renal cell cancer, inhibits both basal and PDGFR $\beta$ -mediated ERK1/2 and AKT activity and decreases cell proliferation in human schwannoma cells, suggesting that this drug constitutes a promising tool to treat schwannomas. We conclude that our schwannoma in vitro model can be used to screen for new therapeutic targets in general and that sorafenib is possible candidate for future clin. trials.
- IT 284461-73-0, Sorafenib
  - RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
    - (BAY 43-9006; growth factor-dependent and growth factor-independent ERK kinase pathway in human schwannoma)
- RN 284461-73-0 CAPLUS
- CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
  - (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

OSC.G 11 THERE ARE 11 CAPLUS RECORDS THAT CITE THIS RECORD (11 CITINGS)
RE.CNT 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN

AN 2007:471132 CAPLUS

DN 147:132413

TI Looking ahead in renal cell carcinoma: integrating new agents in the armamentarium of the urologist

AU Patard, Jean-Jacques

CS Rennes University Hospital, Rennes,

SO European Urology, Supplements (2007), 5(7), 505-509 CODEN: EUSUAU; ISSN: 1569-9056

PB Elsevier B.V.

DT Journal; General Review

LA English

AΒ A review. Urologists play a pivotal role in many aspects of the care of patients with renal cell carcinoma (RCC). However, until recently, in some European countries, they have rarely been involved in the systemic treatment of this disease or in the design of clin. trials. This is undoubtedly set to change with the emergence of new oral, molecularly targeted therapies for RCC. Sorafenib (Nexavar; Bayer Healthcare, West Haven, CT, USA) is one such therapy, which has already been shown to be efficacious and well tolerated for the treatment of RCC. Although targeted agents show great promise for the treatment of RCC, their precise role in the treatment of metastatic disease, and in adjuvant and neoadjuvant settings has yet to be defined. Drawing from their extensive experience of RCC, urologists will be instrumental in the design and application of clin. studies to define the role of targeted therapies in all settings of RCC and, ultimately, to integrate targeted therapies into clin. practice. Through increased understanding of the mol. pathways involved in RCC, research into diagnostic and prognostic markers, and commitment to clin. trials, urologists can be at the forefront of this progress.

IT 475207-59-1, Nexavar

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(mol. targeted therapy with Nexavar was effective and well tolerated in renal cell carcinoma patient)

RN 475207-59-1 CAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 284461-73-0 CMF C21 H16 C1 F3 N4 O3

CM 2

CRN 104-15-4 CMF C7 H8 O3 S

OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
RE.CNT 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
L12 ANSWER 7 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN
     2007:471131 CAPLUS
AN
DN
    147:132412
     New perspectives: an oral multikinase inhibitor in patients with advanced
TI
     RCC
ΑU
     Escudier, Bernard
CS
     Institut Gustave-Roussy, Paris, Fr.
SO
     European Urology, Supplement (2007),
                                           $(7), 499-504
     CODEN: EUSUAU; ISSN: 1569-9036
ΡВ
     Elsevier B.V.
DT
    Journal; General Review
LA
    English
AΒ
               Sorafenib (Nexavar; Bayer Healthcare, West Haven, CT,
    A review.
     USA) is an oral multikinase inhibitor that may provide dual action by
     inhibiting tumor cell proliferation and angiogenesis. Sorafenib was
     recently evaluated in the largest phase 3, randomized trial ever conducted
     in renal cell carcinoma (RCC): Treatment Approaches in Renal Cancer Global
     Evaluation Trial (TARGET). In TARGET, sorafenib significantly increased
     progression-free survival vs. placebo, which led to a change in the study
     protocol allowing patients in the placebo arm of the trial to cross over
     to receive sorafenib. At the time of crossover, sorafenib improved
     overall survival by 39% compared with placebo (hazard ratio = 0.72; 95%
     confidence interval 0.54-0.94; p = 0.02, not significant as per
     O'Brien-Fleming threshold for statistical significance: p = 0.0005).
     Sorafenib continued to show a trend towards improved overall survival at a
     subsequent anal. 6 mo post-crossover. Importantly, 84% of
     sorafenib-treated patients achieved investigator-assessed stable disease
     or better compared with 55% of placebo recipients. Sorafenib was well
     tolerated, had a manageable side-effect profile, and offered benefit with
     no compromise in quality of life. The data from the phase 3 TARGET study
     provided further evidence that sorafenib may be effective in a wide range
     of patients with advanced RCC. Clin. trials are planned to assess the
     potential of sorafenib as combination therapy and in the adjuvant setting.
ΤТ
     475207-59-1, Nexavar
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (oral multikinase inhibitor Nexavar was well tolerated and increased
       progression-free as well as overall survival in patient with advanced
       renal cell carcinoma)
RN
     475207-59-1 CAPLUS
CN
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
     (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-,
     4-methylbenzenesulfonate (1:1) (CA INDEX NAME)
     CM
          1
     CRN 284461-73-0
```

CMF C21 H16 C1 F3 N4 O3

CM 2

CRN 104-15-4 CMF C7 H8 O3 S

OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
L12 ANSWER 8 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN
AN
     2007:471130 CAPLUS
DN
     147:132411
TI
     Adjuvant therapy in renal cell carcinoma: where are we?
ΑU
     Eisen, Tim
CS
     University of Cambridge, Cambridge,
SO
     European Urology, Supplements (2007),
                                           6(7), 492-498
     CODEN: EUSUAU; ISSN: 1569-9058
PΒ
     Elsevier B.V.
    Journal; General Review
DT
LA
    English
AB
     This review summarizes available data and describes planned clin. trials
     designed to evaluate the potential of targeted agents as adjuvant therapy
     for renal cell carcinoma (RCC). Advanced RCC is refractory to standard
     cytotoxic chemotherapy, and clin. trials of adjuvant cytokine therapy in
     this therapeutic setting have not yet demonstrated clear evidence of clin.
     benefit. However, molecularly targeted therapies may offer a new approach
     for adjuvant therapy of this disease. Sorafenib (Nexavar; Bayer
     Healthcare, West Haven, CT, USA) and sunitinib (Sutent; Pfizer Inc, New
     York, NY, USA) are candidates for adjuvant therapy, because they are
     efficacious in the treatment of metastatic RCC and have side-effect
     profiles that can usually be well managed during long-term administration.
     The clin. benefit and tolerability of these agents as adjuvant therapies
     are being investigated in three ongoing phase 3 trials: ASSURE (adjuvant
     sorafenib or sunitinib in unfavorable renal cell carcinoma; Eastern
     Cooperative Oncol. Group 2805), STAR (sunitinib trial in adjuvant renal
     cancer) and SORCE (a phase 3, randomized, double-blind, controlled study
     comparing sorafenib with placebo in patients with resected primary renal
     cell carcinoma at high or intermediate risk of relapse). The results of
     these studies will address important clin. and translational questions,
     the answers to which may help define future treatment strategies and guide
     treatments towards the most appropriate patients.
     475207-59-1, Nexavar
ΙT
     RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological
     activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (adjuvant therapy with Nexavar might be effective in renal cell
        carcinoma patient)
RN
     475207-59-1 CAPLUS
CN
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
     (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-,
     4-methylbenzenesulfonate (1:1) (CA INDEX NAME)
     CM
          1
```

CRN 284461-73-0

CMF C21 H16 C1 F3 N4 O3

Page 21

CM 2

CRN 104-15-4 CMF C7 H8 O3 S

OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)
RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 9 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN

AN 2006:1258840 CAPLUS

DN 146:219796

TI Sorafenib for the treatment of renal cell carcinoma

AU Hughes, Caren L.; Tan, Winston W.; Ferrone, Marcus

CS Oncology Specialty Resident, Division of Pharmacy, MD Anderson Cancer Center, The University of Texas, Houston, TX, USA

SO Journal of Pharmacy Technology (2006), 22(5), 281-288 CODEN: JPTEEB; ISSN: 8755-122

PB Harvey Whitney Books Co.

DT Journal; General Review

LA English

AB A review. Objective: To summarize the pharmacol., development, and clin. application of sorafenib, a specific tyrosine kinase and vascular growth factor inhibitor, for the treatment of renal cell carcinoma (RCC). Data Sources: Clin. literature, including both primary studies and review articles, was obtained by searching MEDLINE (1966-May 2006), using the search terms BAY 43-9006, sorafenib, renal cell carcinoma, and tyrosine kinase inhibitor. Addnl. information was supplied by the manufacturer, Bayer HealthCare Pharmaceuticals. Study Selection and Data Extraction: Review articles, abstrs., and clin. studies related to sorafenib were analyzed. An evaluation of the research exploring sorafenib as a potential therapy for RCC was conducted. Relevant information was then selected and is reviewed in this article. Data Synthesis: Knowledge of the cellular abnormalities that can cause solid tumors has led to the development of medications that block these pathways. Sorafenib is an oral tyrosine kinase inhibitor that both blocks the Raf kinase pathway and inhibits vascular growth factors. Phase I and II trials have demonstrated that sorafenib has activity against RCC. Dermatol. reactions (rash, desquamation), fatique, and hypertension have been the most commonly seen treatment-related adverse events. Sorafenib received FDA approval in Dec. 2005 for treatment of advanced RCC. Conclusions: Soraferib is a novel oral tyrosine kinase inhibitor effective in the treatment of RCC.

IT 284461-73-0, Sorafenib

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(phase I and II trial showed that tyrosine kinase inhibitor sorafenib blocked Raf kinase pathway and inhibited vascular growth factor responsible for angiogenesis and tumor growth in patient with renal cell carcinoma)

RN 284461-73-0 CAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RE.CNT 42 THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 10 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN

AN 2006:1020380 CAPLUS

DN 145:431615

TI Discovery and development of sorafenib: a multikinase inhibitor for treating cancer

AU Wilhelm, Scott; Carter, Christopher; Lynch, Mark; Lowinger, Timothy; Dumas, Jacques; Smith, Roger A.; Schwartz, Brian; Simantov, Ronit; Kelley, Susan

CS Department of Cancer Research, Bayer Pharmaceuticals Corp., West Haven, CT, 06516, USA

SO Nature Reviews Drug Discovery (2006), 5(10), 835-844 CODEN: NRDDAG; ISSN: 1474-1776

PB Nature Publishing Group

DT Journal; General Review

LA English

AB A review. Since the mol. revolution of the 1980s, knowledge of the etiol. of cancer has increased considerably, which has led to the discovery and development of targeted therapies tailored to inhibit cancer-specific pathways. The introduction and refinement of rapid, high-throughput screening technologies over the past decade has greatly facilitated this targeted discovery and development process. Here, the authors describe the discovery and continuing development of sorafenib (previously known as BAY 43-9006), the first oral multikinase inhibitor that targets Raf and affects tumor signaling and the tumor vasculature. The discovery cycle of sorafenib (Nexavar; Bayer Pharmaceuticals) - from initial screening for a lead compound to FDA approval for the treatment of advanced renal cell carcinoma in Dec. 2005 - was completed in just 11 years, with approval being received .apprx.5 years after the initiation of the first Phase I trial.

IT 284461-73-0, Sorafenib

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(discovery and development of sorafenib, a multikinase inhibitor for treating cancer)

RN 284461-73-0 CAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

OSC.G 198 THERE ARE 198 CAPLUS RECORDS THAT CITE THIS RECORD (199 CITINGS)

RE.CNT 93 THERE ARE 93 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 11 OF 11 CAPLUS COPYRIGHT 2011 ACS on STN

AN 2003:736198 CAPLUS

DN 139:301125

TI BAY-43-9006 (Bayer/Onyx)

AU Lee, John T.; McCubrey, James A.

CS Department of Microbiology and Immunology, Brody School of Medicine at East Carolina University, Greenville, NC, 27858-4353, USA

Current Opinion in Investigational Drugs (Thomson Current Drugs) (2003)
4(6), 757-763
CODEN: COIDAZ; ISSN: 1472-4472

PB Thomson Current Drugs

DT Journal; General Review

LA English

AB A review. Bayer and Onyx are developing BAY-43-9006, an oral cytostatic Raf kinase inhibitor for the potential treatment of colorectal and breast cancers, hepatocellular carcinoma and non-small-cell lung cancer, in addition to acute myelogenous leukemia, myelodysplastic syndrome and other cancers. A US IND was filed in May 2000 and by Feb. 2003 BAY-43-9006 was in phase II trials, with phase III trials expected to begin later in 2003.

IT 284461-73-0, BAY 43-9006
RL: ADV (Adverse effect, including toxicity); DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(BAY 43-9006 for treatment of cancer patients)

RN 284461-73-0 CAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

OSC.G 58 THERE ARE 58 CAPLUS RECORDS THAT CITE THIS RECORD (58 CITINGS)
RE.CNT 41 THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
=> s raf or tumor or cancer or carcinoma or adenoma or leukemia
        10045 RAF
        578994 TUMOR
       490811 CANCER
       241830 CARCINOMA
        15739 ADENOMA
       138113 LEUKEMIA
      1042738 RAF OR TUMOR OR CANCER OR CARCINOMA OR ADENOMA OR LEUKEMIA
=> s 110 and 113
L14
    1337 L10 AND L13
=> => d his
     (FILE 'HOME' ENTERED AT 07:44:56 ON 18 JAN 2011)
    FILE 'REGISTRY' ENTERED AT 07:46:14 ON 18 JAN 2011
               STRUCTURE UPLOADED
L1
L2
               STRUCTURE UPLOADED
L3
            43 S L2 SSS SAM
               SCREEN 1840
L4
               SCREEN 2016 OR 2026 OR 1841 OR 2039 OR 2040 OR 2045 OR 20
L5
L6
               STRUCTURE UPLOADED
L7
               OUE L6 AND L4 NOT L5
             4 S L7 SSS SAM
L8
L9
            78 S L7 SSS FUL
    FILE 'CAPLUS' ENTERED AT 07:54:26 ON 18 JAN 2011
L10
          1515 S L9
         10038 S BAYER?
L11
L12
            11 S L10 AND L11
L13
      1042738 S RAF OR TUMOR OR CANCER OR CARCINOMA OR ADENOMA OR LEUKEMIA
L14
          1337 S L10 AND L13
    FILE 'USPATFULL' ENTERED AT 07:58:05 ON 18 JAN 2011
=> s 19
L15
         335 L9
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=> d 115 301-335 bib, ab, hitstr

Page 27

L15 ANSWER 301 OF 335 USPATFULL on STN 2005:247130 USPATFULL ΑN Compositions and methods to increase the effect of a neurotoxin ΤТ treatment INDavid, Mathaniel E., San Francisco, CA, UNITED STATES VVII NewCo 2008, Inc., Menlo Park, CA, UNITED STATES (U.S. corporation) PAPΙ US 20050214325 A1 / 20050929 ΑI US 2004-810391 A1 \ 20040326 /(10) DT Utility FS APPLICATION LREP WILSON SONSINI GOODRICH & ROSATI, 650 PAGE MILL ROAD, PALO ALTO, CA, 94304-1050, US CLMN Number of Claims: 40 ECL Exemplary Claim: 1 1 Drawing Page(s) DRWN LN.CNT 1120 CAS INDEXING IS AVAILABLE FOR THIS PATENT. The present invention discloses compositions and methods for enhancing AB the effect (e.g., duration) of a neurotoxin treatment. The compositions herein include neurotoxins and neuron growth inhibitors. Such compositions are administered locally to treat or prevent conditions, such as dermatological conditions, urological conditions, thyroid conditions, optical conditions, and neurological conditions. 284461-73-0, BAY-43-9006 ΙT (compns. and methods to increase effect of neurotoxin treatment) RN 284461-73-0 USPATFULL CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

L15 ANSWER 302 OF 335 USPATFULL on STN

AN 2005:183990 USPATFULL

TI JAK/STAT inhibitors and MAPK/ERK inhibitors for RSV infection

IN Mohapatra, Shyam S., Tampa, UNITED STATES

PI US 20050159385 AZ 20050721 AI US 2004-18954 71 20041220 (11) PRAI US 2003-531052P 20031219 (60)

DT Utility FS APPLICATION

LREP SALIWANCHIK LLOYD & SALIWANCHIK, A PROFESSIONAL ASSOCIATION, PO BOX 142950, GAINESVILLE, FL, 32614-2950, US

CLMN Number of Claims: 20 ECL Exemplary Claim: 1 DRWN 17 Drawing Page(s) LN.CNT 2773

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention concerns a method for treating or reducing the likelihood of developing a respiratory syncytial virus (RSV) infection in a subject by administering an effective amount of an inhibitor of the janus kinase (JAK)/signal transducer and activator of transcription (STAT) signaling pathway or the mitogen-activated kinase (MAPK)/extracellular signal-regulated kinase (ERK1/2) signaling pathway to the subject. Another aspect of the invention concerns a pharmaceutical composition that includes an inhibitor of JAK/STAT or MAPK/ERK signaling to the subject; and a pharmaceutically acceptable carrier. Another aspect of the invention concerns a method for identifying agents useful for treating or reducing the likelihood of developing an RSV infection

IT 284461-73-0, BAY 43-9006

(JAK/STAT inhibitors and MAPK/ERK inhibitors for respiratory syncytial virus infection treatment)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CAINDEX NAME)

L15 ANSWER 303 OF 335 USPATFULL on STN

AN 2005:171786 USPATFULL

TI IAP nucleobase oligomers and oligomeric complexes and uses thereof

IN LaCasse, Eric, Ottawa, CANADA

McManus, Daniel, Ottawa, CANADA

PI US 20050148535 A1 20050707

AI US 2004-975974 A1 20041028 (10) PRAI US 2003-516192P 20031030 (60)

DT Utility

FS APPLICATION

LREP CLARK & ELBING LLP, 101 FEDERAL STREET, BOSTON, MA, 02110, US

CLMN Number of Claims: 48 ECL Exemplary Claim: 1

DRWN 15 Drawing Page(s)

LN.CNT 3022

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides nucleobase oligomers and oligomer complexes that inhibit expression of an IAP polypeptide, and methods for using them to induce apoptosis in a cell. The nucleobase oligomers and oligomer complexes of the present invention may also be used to form pharmaceutical compositions. The invention also features methods for enhancing apoptosis in a cell by administering a nucleobase oligomer or oligomer complex of the invention in combination with a chemotherapeutic or chemosensitizing agent.

IT 284461-73-0, BAY-43-9006

(human protein IAP (inhibitor of apoptosis protein) nucleobase oligomers, including dsRNA, shRNA, and siRNA, and their use for enhancing apoptosis in cancer therapy)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

L15 ANSWER 304 OF 335 USPATFULL on STN

AN 2005:138567 USPATFULL

TI Methods and reagents for the treatment of proliferative diseases

IN LaCasse, Eric, Ottawa, CANADA McManus, Daniel, Ottawa, CANADA

Durkin, Jon P., Montreal, CANADA

PI US 20050119217 A1 20050602

AI US 2004-975790 A1 20041028 (10) PRAI US 2003-516263P 20031030 (60)

DT Utility

FS APPLICATION

LREP CLARK & ELBING LLP, 101 FEDERAL STREET, BOSTON, MA, 02110, US

CLMN Number of Claims: 58
ECL Exemplary Claim: 1
DRWN 34 Drawing Page(s)

LN.CNT 5896

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention features methods, compositions, and kits for treating a patient having a proliferative disease.

IT 284461-73-0, BAY-43-9006

(sequences of antisense IAP (inhibitor of apoptosis protein) oligomers and their use for treatment of proliferative diseases with chemotherapeutic agent)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CAINDEX NAME)

L15 ANSWER 305 OF 335 USPATFULL on STN

AN 2005:137954 USPATFULL

TI Method for selecting drug sensitivity-determining factors and method for predicting drug sensitivity using the selected factors

IN Aoki, Yuko, Kanagawa, JAPAN

Hasegawa, Kiyoshi, Kanagawa, JAPAN

Ishii, Nobuya, Kanagawa, JAPAN

Mori, Kazushige, Kanagawa, JAPAN

PI US 20050118600 A1 20050602

AI US 2003-507389 A1 20020313 (10)

WO 2002-JP2354 20020313

DT Utility

FS APPLICATION

LREP FISH & RICHARDSON PC, 225 FRANKLIN ST, BOSTON, MA, 02110, US

CLMN Number of Claims: 31

ECL Exemplary Claim: 1

DRWN 7 Drawing Page(s)

LN.CNT 2028

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AΒ Based on drug sensitivity data and extensive gene expression data, a model was constructed by multivariate analysis with the partial least squares method type 1. Further, the model was optimized using modeling power and genetic algorithm. Thereby, the degree of contribution of the respective genes to drug sensitivity was determined to select genes with a high degree of contribution. In addition, the levels of gene expression in specimens were analyzed, and then the drug sensitivity was predicted based on the model. The predicted values agreed well with those drug sensitivity values determined experimentally. The drug sensitivity-predicting method provided by the present invention enables assessment of the effectiveness of a drug prior to administration using small quantities of specimens associated with diseases such as cancer. Since this enables the selection of the most suitable drug for each patient, the present invention is very useful in improving a patient's quality of life (QOL).

IT 284461-73-0, BAY 439006

(method for selecting antitumor drug sensitivity-determining factors and predicting antitumor drug sensitivity using the selected factors)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

```
L15 ANSWER 306 OF 335 USPATFULL on STN
       2005:69562 USPATFULL
ΑN
ΤТ
       Diaryl ureas for diseases mediated by PDGFR
       Wilhelm, Scott, Orange, CT, UNITED STATES
Dumas, Jacques, Bethany, CT, UNITED STATES
ΙN
                                                           common inventor
       Ladouceur, Gaetan, Guilford, CT, UNITED STATES
       Lynch, Mark, Madison, CT, UNITED STATES
       Scott, William, Guilford, CT, UNITED STATES
PΙ
       US 20050059703
                           A1 20050317
       US 2004-848567
                           A1 20040519 (10)
ΑТ
PRAI
       US 2004-556062P
                                20040325 (60)
       US 2003-520399P
                                20031117 (60)
       US 2003-471735P
                                20030520 (60)
DT
       Utility
FS
       APPLICATION
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
LREP
       1400, ARLINGTON, VA, 22201
       Number of Claims: 74
CLMN
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 1901
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention provides methods for treating and/or preventing
       conditions and diseases in humans and other mammals that are associated
       with and/or mediated by signal transduction pathways comprising
       platelet-derived growth factor receptor (PDGFR) by administering diaryl
       ureas of Formula I. The present invention also provides devices and
       methods for treating, ameliorating, preventing, or modulating restenosis
       following angioplastic surgery or other invasive procedures that affect
       or injure the vascular system, and graft rejection following
       transplantation of a donor tissue into a host, where a stent or other
       omplantable device comprises an effective amount of diaryl ureas of
       Formula I.
    284461-73-0 284461-74-1 284462-18-6
      475207-59-1 583840-03-3 583840-04-4
        (diaryl ureas for prevention and/or treatment of diseases mediated by
        platelet-derived growth factor receptor)
RN
     284461-73-0 USPATFULL
CN
     2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```

RN 284461-74-1 USPATFULL CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & \\ & & & \\ & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 475207-59-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME)

CM 1

CRN 284461-73-0

CMF C21 H16 C1 F3 N4 O3

CM 2

CRN 104-15-4 CMF C7 H8 O3 S

RN 583840-03-3 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 1-oxide (CA INDEX NAME)

RN 583840-04-4 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-, 1-oxide (CA INDEX NAME)

$$H_2N-C$$
 $O$ 
 $NH-C-NH$ 
 $CF_3$ 
 $C1$ 

```
L15 ANSWER 307 OF 335 USPATFULL on STN
       2005:69531 USPATFULL
ΑN
       Novel farnesyl protein transferase inhibitors as antitumor agents
ΤТ
ΙN
       Zhu, Hugh Y., Scotch Plains, NJ, UNITED STATES
       Cooper, Alan B., West Caldwell, NJ, UNITED STATES
       Desai, Jagdish A., Monroe Township, NJ, UNITED STATES
       Wang, James J-S, Westfield, NJ, UNITED STATES
       Rane, Dinanath F., Morganville, NJ, UNITED STATES
       Doll, Ronald J., Convent Station, NJ, UNITED STATES
       Njoroge, F. George, Warren, NJ, UNITED STATES
       Girijavallabhan, Viyyoor M., Parsipanny, NJ, UNITED STATES
PΑ
       SCHERING CORPORATION (U.S. corporation)
       US 20050059672
PΙ
                           A1 20050317
       US 7557107
                               20090707
                           В2
       US 2004-911340
                               Z0040804 (10)
ΑТ
                           A 1
       US 2003-493269P
                               20030807
                                         §60)
PRAI
       US 2003-498509P
                                20030828
                                         ((60)
DT
       Utility
FS
       APPLICATION
LREP
       SCHERING-PLOUGH CORPORATION, PATENT DEPARTMENT (K-6-1, 1990), 2000
       GALLOPING HILL ROAD, KENILWORTH, NJ, 07033-0530
CLMN
       Number of Claims: 120
       Exemplary Claim: 1
ECL
       No Drawings
DRWN
LN.CNT 4090
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Disclosed are novel tricyclic compounds of the formula:
       and a pharmaceutically acceptable salts or solvates thereof. The
       disclosed are pharmaceutical compositions comprising the compounds of
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compounds are useful for inhibiting farnesyl protein transferase. Also formula (I). Also disclosed are uses of the compounds of formula (I) for the manufacture of a medicament for the treatment of cancer.

ΤT 284461-73-0, Bay 43-9006

(coadministration; preparation of piperazinylbenzocycloheptapyridines as farnesyl protein transferase inhibitors useful as antitumor agents)

RN 284461-73-0 USPATFULL

2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-CN

> (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

L15 ANSWER 308 OF 335 USPATFULL on STN 2005:56618 USPATFULL ΑN BRAF mutation T1796A in thyroid cancers ТΤ ΙN Sidransky, David, Baltimore, MD, UNITED STATES Cohen, Yoram, Baltimore, MD, UNITED STATES Zhao, Ming, Clarksville, MD, UNITED STATES The Johns Hopkins University, Baltimore, MD, UNITED STATES, 21218 (U.S. PAcorporation) US 20050048533 A1 20050303 PΙ US 7378233 B2 20080527 US 2004-821203 A1 / 20040409 (10) AΙ PRAI 20030412/(60) US 2003-462046P DT Utility APPLICATION FS BANNER & WITCOFF, 1001 G STREET N W, SUITE 1100, WASHINGTON, DC, 20001 LREP Number of Claims: 24 CLMN Exemplary Claim: 1 ECL DRWN 2 Drawing Page(s)

LN.CNT 1021

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The BRAF gene has been found to be activated by mutation in human cancers, predominantly in malignant melanoma. We tested 476 primary tumors, including 214 lung, 126 head and neck, 54 thyroid, 27 bladder, 38 cervical, and 17 prostate cancers, for the BRAF T1796A mutation by polymerase chain reaction (PCR)-restriction enzyme analysis of BRAF exon 15. In 24 (69%) of the 35 papillary thyroid carcinomas examined, we found a missense thymine (T)-adenine (A) transversion at nucleotide 1796 in the BRAF gene (T1796A). The T1796A mutation was detected in four lung cancers and in six head and neck cancers but not in bladder, cervical, or prostate cancers. Our data suggest that activating BRAF mutations may be an important event in the development of papillary thyroid cancer. Moreover, BRAF mutation reliably predicts a poor prognosis for papillary thyroid carcinomas.

IT 284461-73-0, BAY 43-9006

(detection of BRAF transversion mutation for diagnosis of malignant thyroid cancer and uses of Ras-Raf-MAPK or Raf/MEK/ERK signaling pathway inhibitor in treating thyroid cancer)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

 $\label{lem:carbonyl} $$ (trifluoromethyl) phenyl] amino] carbonyl] amino] phenoxy]-N-methyl- (CA INDEX NAME)$ 

```
L15 ANSWER 309 OF 335 USPATFULL on STN
ΑN
       2005:44298 USPATFULL
       Nove (bicyclic) urea derivatives useful in the treatment of cancer and
ΤТ
       other disorders
      Qumas, Jacques Bethany, CT, UNITED STATES
ΙN
       Boyer, Stephen, Fairfield, CT, UNITED STATES
       Verma, Sharad, New Haven, CT, UNITED STATES
                                                         Claims contain a bicyclic group
       Adnane, Lila, Madison, CT, UNITED STATES
       Chen, Yuanwei, North Haven, CT, UNITED STATES
       Lee, Wendy, Hamden, CT, UNITED STATES
       Phillips, Barton, New Haven, CT, UNITED STATES
       Smith, Roger A., Madison, CT, UNITED STATES
       Scott, William J., Guildford, CT, UNITED STATES
       Burke, Jennifer, New Haven, CT, UNITED STATES
       Chen, Jianqing, New Haven, CT, UNITED STATES
       Chen, Zhi, Hamden, CT, UNITED STATES
       Fan, Jianmei, Hamden, CT, UNITED STATES
       Miranda, Karl, North Haven, CT, UNITED STATES
       Raudenbush, Brian, Charlton, MA, UNITED STATES
       Redman, Aniko, Derby, CT, UNITED STATES
       Shao, Jianxing, Acton, MA, UNITED STATES
       Su, Ning, Hamden, CT, UNITED STATES
       Wang, Gan, Wallingford, CT, UNITED STATES
       Yi, Lin, Milford, CT, UNITED STATES
       Zhu, Qingming, West Haven, CT, UNITED STATES
PΙ
       US 20050038031 A1 20050217
ΑI
       US 2004-788426
                           A1 20040301 (10)
PRAI
       US 2003-450323P
                               20030228 (60)
       US 2003-450324P
                               20030228 (60)
DT
       Utility
FS
       APPLICATION
LREP
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
CLMN
       Number of Claims: 30
ECL
       Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 4157
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AΒ
       This invention relates to novel diaryl ureas, pharmaceutical
       compositions containing such compounds and the use of those compounds or
       compositions for treating hyper-proliferative and angiogenesis
       disorders, as a sole agent or in combination with cytotoxic therapies.
    284461-73-0, Bay 43-9006
ΤT
        (coadministration; preparation of ureidophenoxycyanopyridines as anticancer
        drugs)
     284461-73-0 USPATFULL
RN
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
CN
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```

```
L15 ANSWER 310 OF 335 USPATFULL on STN
       2005:38118 USPATFULL
ΑN
       2-0xo(1,3,5-perhydrotriazapine derivatives useful in the treatment of
ΤТ
       hyper-proliferative, angiogenesis, and inflammatory disorders
ΙN
       Boyer, Stophen, Fairfield, CT, UNITED STATES
      (Dumas, Jacques) Bethany, CT, UNITED STATES
       Phillips, Barton, New Haven, CT, UNITED STATES
       Scott, William J., Guildford, CT, UNITED STATES
       Smith, Roger A., Madison, CT, UNITED STATES
       Chen, Jianqing, New Haven, CT, UNITED STATES
       Jones, Benjamin, Hamden, CT, UNITED STATES
       Wang, Gan, Wallingford, CT, UNITED STATES
                          A1 20050210
PΙ
       US 20050032798
                                                        no ODP
       US 2004-788405
                           A1 20040301 (10)
ΑI
       US 2003-450323P
PRAI
                               20030228 (60)
       US 2003-450324P
                               20030228 (60)
       US 2003-450348P
                               20030228 (60)
DT
       Utility
FS
       APPLICATION
LREP
      MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
CLMN
      Number of Claims: 46
       Exemplary Claim: 1
ECL
      No Drawings
DRWN
LN.CNT 2600
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to novel diaryl ureas, pharmaceutical
       compositions containing such compounds and the use of those compounds or
       compositions for treating hyper-proliferative and angiogenesis
       disorders, as a sole agent or in combination with cytotoxic therapies.
TT
    284461-73-0, Bay 43-9006
        (coadministration; preparation of ureidophenoxycyanopyridines as anticancer
        drugs)
RN
     284461-73-0 USPATFULL
CN
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```

```
L15 ANSWER 311 OF 335 USPATFULL on STN
       2004:299960 USPATFULL
ΑN
ΤТ
       Novel (cyanopyridine) derivatives useful in the treatment of cancer and
       other disorders
       Scott, William J., Guilford, CT, UNITED STATES
ΙN
     (Dumas, Jacques, Bethany, CT, UNITED STATES
      Boyer, Stephen, Hilden, GERMANY, FEDERAL REPUBLIC OF
       Lee, Wendy, Hamden, CT, UNITED STATES
       Chen, Yuanwei, North Haven, CT, UNITED STATES
       Phillips, Barton, New Haven, CT, UNITED STATES
       Verma, Sharad, New Haven, CT, UNITED STATES
       Chen, Jianqing, New Haven, CT, UNITED STATES
       Chen, Zhi, Hamden, CT, UNITED STATES
       Fan, Jianmei, Hamden, CT, UNITED STATES
       Raudenbush, Brian, Charlton, MA, UNITED STATES
       Redman, Aniko, Derby, CT, UNITED STATES
       Yi, Lin, Milford, CT, UNITED STATES
       Zhu, Qingming, West Haven, CT, UNITED STATES
       Adnane, Lila, Madison, CT, UNITED STATES
PΙ
       US 20040235829
                          A1 20041125
       US 7557129
                           B2 20090707
                                                no ODP
                           A1 20040227 (10)
ΑI
      US 2004-788029
                               20030228 (60)
PRAI
      US 2003-450323P
                               20030228 (60)
       US 2003-450324P
       US 2003-450348P
                               20030228 (60)
DT
      Utility
FS
      APPLICATION
LREP
      MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
      Number of Claims: 63
CLMN
      Exemplary Claim: 1
ECL
DRWN
      No Drawings
LN.CNT 2828
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AΒ
       This invention relates to novel diaryl ureas, pharmaceutical
       compositions containing such compounds and the use of those compounds or
       compositions for treating hyper-proliferative and angiogenesis
       disorders, as a sole agent or in combination with cytotoxic therapies.
    284461-73-0, Bay 43-9006
IΤ
        (coadministration; preparation of ureidophenoxycyanopyridines as anticancer
        drugs)
RN
     284461-73-0 USPATFULL
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
CN
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```

```
L15 ANSWER 312 OF 335 USPATFULL on STN
ΑN
       2004:292848 USPATFULL
ΤТ
       Substituted pyridine derivatives useful in the treatment of cancer and
       Qther-drsorders
      Qumas, Jacques Bethany, CT, UNITED STATES
ΙN
       Lee, Wendy, Hamden, CT, UNITED STATES
       Chen, Yuanwei, North Haven, CT, UNITED STATES
       Adnane, Lila, Madison, CT, UNITED STATES
       Scott, William J., Guilford, CT, UNITED STATES
       Verma, Sharad, New Haven, CT, UNITED STATES
       Chen, Jianqing, New Haven, CT, UNITED STATES
       Chen, Zhi, Hamden, CT, UNITED STATES
       Yi, Lin, Milford, CT, UNITED STATES
PΙ
       US 20040229937
                          A1 20041118
                                               ABN
       US 2004-789446
ΑI
                           A1 20040301 (10)
       US 2003-450323P
                               20030228 (60)
PRAI
       US 2003-450324P
                               20030228 (60)
       US 2003-450348P
                               20030228 (60)
DT
       Utility
FS
       APPLICATION
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
LREP
       1400, ARLINGTON, VA, 22201
       Number of Claims: 25
CLMN
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 2564
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB
       This invention relates to novel diaryl ureas, pharmaceutical
       compositions containing such compounds and the use of those compounds or
       compositions for treating hyper-proliferative and angiogenesis
       disorders, as a sole agent or in combination with cytotoxic therapies.
    284461-73-0, Bay 43-9006
ΙT
        (coadministration; preparation of ureidophenoxycyanopyridines as anticancer
        drugs)
RN
     284461-73-0 USPATFULL
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
CN
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```

L15 ANSWER 313 OF 335 USPATFULL on STN

AN 2004:165963 USPATFULL

TI Method for treating diseases associated with abnormal kinase activity

IN Lyons, John, Moraga, CA, UNITED STATES

Rubinfeld, Joseph, Danville, CA, UNITED STATES

PI US 20040127453 A1 20040701

US 6998391 B2 20060214

AI US 2002-206854 A1 20020726 (10)

RLI Continuation-in-part of Ser. No. US 2002-71849, filed on 7 Feb 2002,

PENDING

DT Utility

FS APPLICATION

LREP WILSON SONSINI GOODRICH & ROSATI, 650 PAGE MILL ROAD, PALO ALTO, CA,

943041050

CLMN Number of Claims: 66

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 1941

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AΒ Methods are provided for treating diseases associated with abnormal activity of kinases. The method comprises: administering a DNA methylation inhibitor to the patient in therapeutically effective amount; and administering a kinase inhibitor to the patient in therapeutically effective amount, such that the in vivo activity of the kinase is reduced relative to that prior to the treatment. The method can be used to treat cancer associated with abnormal activity of kinases such as phosphatidylinositol 3'-kinase (PI3K), protein kinases including serine/threonine kinases such as Raf kinases, protein kinase kinases such as MEK, and tyrosine kinases such as those in the epidermal growth factor receptor family (EGFR), platelet-derived growth factor receptor family (PDGFR), vascular endothelial growth factor receptor (VEGFR) family, nerve growth factor receptor family (NGFR), fibroblast growth factor receptor family (FGFR) insulin receptor family, ephrin receptor family, Met family, Ror family, c-kit family, Src family, Fes family, JAK family, Fak family, Btk family, Syk/ZAP-70 family, and Abl family.

IT 284461-73-0, BAY 43-9006

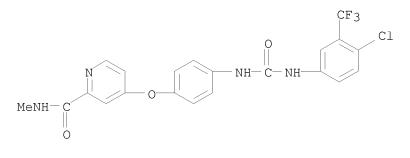
(Raf kinase inhibitor; treating diseases associated with abnormal tyrosine kinase activity by administering DNA methylation inhibitors and tyrosine kinase inhibitors)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

 $\label{lem:carbonyl} $$ (trifluoromethyl) phenyl] amino] carbonyl] amino] phenoxy]-N-methyl- (CAINDEX NAME)$ 

L15 ANSWER 314 OF 335 USPATFULL on STN 2003:330550 USPATFULL ΑN Aryl urea compounds in combination with other cytostatic or cytotoxic ΤТ agents for treating human cancers IN Carter, Christopher A., Guilford, CT, UNITED STATES Gibson, Neil, East Northport, NY, UNITED STATES Hibner, Barbara, Madison, CT, UNITED STATES Humphrey, Rachel W., Woodbridge, CT, UNITED STATES Trail, Pamela, Madison, CT, UNITED STATES Vincent, Patrick W., Cheshire, CT, UNITED STATES Zhai, Yifan, Guilford, CT, UNITED STATES PABAYER CORPORATION, Pittsburgh, PA, UNITED STATES (U.S. corporation) PΙ US 20030232765 A1 20031218 ABN US 2002-308187 A1 20021203 (10) ΑI US 2001-334609P 20011203 (60) PRAI DТ Utility APPLICATION FS MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE LREP 1400, ARLINGTON, VA, 22201 CLMN Number of Claims: 9 ECL Exemplary Claim: 1 DRWN 5 Drawing Page(s) LN.CNT 1005 CAS INDEXING IS AVAILABLE FOR THIS PATENT. This invention relates to aryl urea compounds in combination with cytotoxic or cytostatic agents for use in treating raf kinase mediated diseases such as cancer. 475207-59-1 ΤT (aryl urea compds. in combination with other cytostatic or cytotoxic agents for treating human cancers and other raf kinase-mediated diseases) 475207-59-1 USPATFULL RN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-CN (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 4-methylbenzenesulfonate (1:1) (CA INDEX NAME) CM 1 CRN 284461-73-0 CMF C21 H16 C1 F3 N4 O3

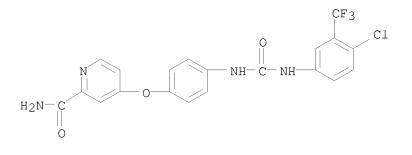


CM 2

CRN 104-15-4

CMF C7 H8 O3 S

L15 ANSWER 315 OF 335 USPATFULL on STN ΑN 2003:307010 USPATFULL Aryl ureas as kinase inhibitors ΤТ Dumas, Jacques, Orange, CT, UNITED STATES ΙN Scott, William J., Guilford, CT, UNITED STATES Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF Chien, Du-Shieng, Guilford, CT, UNITED STATES Nassar, Ala, Milford, CT, UNITED STATES Lee, Wendy, Hamden, CT, UNITED STATES Bjorge, Susan, Milford, CT, UNITED STATES Musza, Laszlo L., Guilford, CT, UNITED STATES PΑ BAYER CORPORATION, Pittsburgh, PA, UNITED STATES (U.S. corporation) PΙ US 20030216446 A1 20031120 US 2003-361859 ΑI Α1 20030211 (10) now allowed US 2002-354937P 20020211 (60) PRAI N-oxide compounds DТ Utility APPLICATION FS MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE LREP 1400, ARLINGTON, VA, 22201 CLMN Number of Claims: 73 ECL Exemplary Claim: 1 DRWN No Drawings LN.CNT 1856 CAS INDEXING IS AVAILABLE FOR THIS PATENT. This invention relates to new aryl ureas and methods for their synthesis. The inventive compounds are useful in the treatment of (i) raf mediated diseases, for example, cancer, (ii) p38 mediated diseases such as inflammation and osteoporosis, and (iii) VEGF mediated diseases such as angiogenesis disorders. 284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-[2carbamoyl(4-pyridyloxy)phenyl]urea 284462-18-6P 583840-03-3P 583840-04-4P 583840-09-9P (preparation of aryl ureas for therapeutic use as kinase inhibitors) RN 284461-74-1 USPATFULL 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-CN (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)



RN 284462-18-6 USPATFULL
CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-(CA INDEX NAME)

RN 583840-03-3 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 1-oxide (CA INDEX NAME)

RN 583840-04-4 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-, 1-oxide (CA INDEX NAME)

RN 583840-09-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

IT 583840-05-5P 583840-06-6P 583840-07-7P

583840-08-8P

(preparation of aryl ureas for therapeutic use as kinase inhibitors)

RN 583840-05-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(hydroxymethyl), 1-oxide (CA INDEX NAME)

RN 583840-06-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 1-oxide (CA INDEX NAME)

RN 583840-07-7 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(hydroxymethyl), 1-oxide (CA INDEX NAME)

RN 583840-08-8 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-, 1-oxide (CA INDEX NAME)

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

L15 ANSWER 316 OF 335 USPATFULL on STN

AN 2003:306960 USPATFULL

TI Pyridine, quinoline, and isoquinoline N-oxides as kinase inhibitors

IN Dumas, Jacques, Bethany, CT, UNITED STATES

Scott, William J., Guilford, CT, UNITED STATES

Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF

PA BAYER CORPORATION, Pittsburgh, PA (U.S. corporation)

PI US 20030216396 A1 20031120

AI US 2003-361850 A1 20030211 (10)

PRAI US 2002-354935P 20020211 (60)

DT Utility

FS APPLICATION

LREP MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE 1400, ARLINGTON, VA, 22201

CLMN Number of Claims: 35

ECL Exemplary Claim: 1

DRWN No Drawings

LN.CNT 2076

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention relates to urea compounds containing a pyridine, quinoline, or isoquinoline functionality which is oxidized at the nitrogen heteroatom and which are useful in the treatment of (i) raf mediated diseases, for example, cancer, (ii) p38 mediated diseases such as inflammation and osteoporosis, and (iii) VEGF mediated diseases such as angiogenesis disorders.

IT 284461-73-0

(preparation of aryl ureas containing pyridine, quinoline and isoquinoline N-oxide functionality as kinase inhibitors)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

IT 284461-74-1P

(preparation of aryl ureas containing pyridine, quinoline and isoquinoline N-oxide functionality as kinase inhibitors)

RN 284461-74-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

IT 583840-03-3P 583840-04-4P

(preparation of aryl ureas containing pyridine, quinoline and isoquinoline  $\mathbb{N}$ -oxide functionality as kinase inhibitors)

RN 583840-03-3 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-, 1-oxide (CA INDEX NAME)

RN 583840-04-4 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-, 1-oxide (CA INDEX NAME)

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L15 ANSWER 317 OF 335 USPATFULL on STN
       2003:294854 USPATFULL
AN
       OMEGA-CARBOXYARYL SUBSTITUTED DIPHENYL UREAS AS RAF KINASE INHIBITORS
ΤТ
ΙN
       Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF
       Dumas, Jacques, Orange, CT, UNITED STATES
       Khire, Uday, Hamden, CT, UNITED STATES
       Lowinger, Timothy B., Nishinomiya City, JAPAN
       Scott, William J., Guilford, CT, UNITED STATES
       Smith, Roger A., Madison, CT, UNITED STATES
       Wood, Jill E., Hamden, CT, UNITED STATES
       Monahan, Mary-Katherine, Hamden, CT, UNITED STATES
       Natero, Reina, Hamden, CT, UNITED STATES
       Renick, Joel, Milford, CT, UNITED STATES
       Sibley, Robert N., North Haven, CT, UNITED STATES
       BAYER CORPORATION, Pittsburgh, PA (non-U.S. corporation)
PA
PТ
       US 20030207872
                           A1 20031106
                                              ABN
ΑI
       US 2002-42226
                           A1
                               20020111 (10)
DT
       Utility
FS
       APPLICATION
LREP
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
CLMN
       Number of Claims: 67
       Exemplary Claim: 1
ECL
       No Drawings
DRWN
LN.CNT 3713
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to the use of a group of aryl ureas in treating
       raf mediated diseases, and pharmaceutical compositions for use in such
       therapy.
   284461-73-0P 284461-74-1P 284461-82-1P
      284461-88-7P 284462-04-0P 284462-05-1P
      284462-17-5P 284462-18-6P 284462-21-1P
      604813-04-9P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-[[3-[5-
      (2-dimethylaminoethyl)carbamoyl]pyridyl]oxy]phenyl]urea
        (preparation of ω-carboxyaryl substituted di-Ph ureas as raf kinase
        inhibitors for treating raf-mediated diseases such as cancerous cell
        growth)
RN
     284461-73-0 USPATFULL
CN
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
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```

RN 284461-74-1 USPATFULL
CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & \\ & & & \\ & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-05-1 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-(CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

RN 604813-04-9 USPATFULL

CN 3-Pyridinecarboxamide, 6-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

L15 ANSWER 318 OF 335 USPATFULL on STN ΑN 2003:294852 USPATFULL Aryl wreas with angiogenisis inhibiting activity ΤТ ΙN (Dumas, Jacques, Orange, CT, UNITED STATES Scott, William J., Guilford, CT, UNITED STATES Elting, James, Madison, CT, UNITED STATES Hatoum-Makdad, Holia, Hamden, CT, UNITED STATES BAYER CORPORATION, Pittsburgh, PA (U.S. corporation) PAPΙ US 20030207870 A1 20031106 US 7838541 B2 20101123 retinopathy ΑI US 2003-361858 A1 20030211 (10) **VEGF** PRAI US 2002-354950P 20020211 (60) DT Utility FS APPLICATION MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE LREP 1400, ARLINGTON, VA, 22201 Number of Claims: 32 CLMN ECL Exemplary Claim: 1 DRWN No Drawings LN.CNT 2356 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB This invention relates to methods of using aryl ureas to treat diseases mediated by the VEGF induced signal transduction pathway characterized by abnormal angiogenesis or hyperpermeability processes. 284461-73-0P 284461-74-1P ΙT (preparation of aryl ureas with angiogenesis inhibiting activity) RN 284461-73-0 USPATFULL CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-74-1 USPATFULL
CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & &$$

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L15 ANSWER 319 OF 335 USPATFULL on STN
       2003:271082 USPATFULL
AΝ
       Antibodies that immunospecifically bind to trail receptors
TΤ
ΙN
       Salcedo, Theodora, Montgomery Village, MD, UNITED STATES
       Ruben, Steven M., Brookeville, MD, UNITED STATES
       Rosen, Craig A., Laytonsville, MD, UNITED STATES
       Albert, Vivian R., Rockville, MD, UNITED STATES
       Dobson, Claire, Cambridge, UNITED KINGDOM
       Vaughan, Tristan, Cambridge, UNITED KINGDOM
      Human Genome Sciences Inc., Rockville, MD, UNITED STATES, 20850 (U.S.
PA
       Corporation)
                          A1 20031009
PΙ
       US 20030190685
                          B2 20060620
       US 7064189
       US 2002-139785
                          A1 20020507 (10)
ΑТ
       US 2001-293473P
                               20010525 (60)
PRAI
       US 2001-294981P
                               20010604 (60)
       US 2001-309176P
                               20010802 (60)
       US 2001-323807P
                               20010921 (60)
       US 2001-327364P
                               20011009 (60)
                               20011107 (60)
       US 2001-331044P
                               20011114 (60)
       US 2001-331310P
       US 2001-341237P
                               20011220 (60)
       US 2002-369860P
                               20020405 (60)
DΤ
       Utility
      APPLICATION
FS
LREP
      HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, ROCKVILLE, MD, 20850
CLMN
      Number of Claims: 77
ECL
       Exemplary Claim: 1
DRWN
       3 Drawing Page(s)
LN.CNT 11875
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AΒ
       The present invention relates to antibodies and related molecules that
       immunospecifically bind to TRAIL receptor, TR4. Such antibodies have
       uses, for example, in the prevention and treatment of cancers and other
       proliferative disorders. The invention also relates to nucleic acid
      molecules encoding anti-TR4 antibodies, vectors and host cells
       containing these nucleic acids, and methods for producing the same. The
      present invention relates to methods and compositions for preventing,
       detecting, diagnosing, treating or ameliorating a disease or disorder,
       especially cancer and other hyperproliferative disorders, comprising
       administering to an animal, preferably a human, an effective amount of
       one or more antibodies or fragments or variants thereof, or related
       molecules, that immunospecifically bind to TRAIL receptor TR4.
    284461-73-0, BAY 43-9006
ΤТ
        (anti-human TRAIL receptor TR4 antibodies and scFvs for diagnosis and
        treatment of cancer or hyperproliferative disease)
     284461-73-0 USPATFULL
RN
     2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
CN
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
```

INDEX NAME)

```
L15 ANSWER 320 OF 335 USPATFULL on STN
       2003:258389 USPATFULL
AN
ΤТ
       omega-carboxyaryl substituted diphenyl ureas as raf kinase inhibitors
ΙN
       Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF
       Dumas, Jacques, Orange, CT, UNITED STATES
       Khire, Uday, Hamden, CT, UNITED STATES
       Lowinger, Timothy B., Nishinomiya City, JAPAN
       Scott, William J., Guilford, CT, UNITED STATES
       Smith, Roger A., Madison, CT, UNITED STATES
                                                               Applicant's
       Wood, Jill E., North Haven, CT, UNITED STATES
       Monahan, Mary-Katherine, Hamden, CT, UNITED STATES
       Natero, Reina, Hamden, CT, UNITED STATES
       Renick, Joel, San Diego, CA, UNITED STATES
       Sibley, Robert N., North Haven, CT, UNITED STATES
       BAYER CORPORATION, Piittsburgh, PA (non-U.S. corporation)
PA
PТ
       US-20030181442
                               20030925
                           A1
ΑI
       US 2001-993647
                               20011127 (9)
                           Α1
DT
       Utility
FS
       APPLICATION
LREP
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
CLMN
       Number of Claims: 67
       Exemplary Claim: 1
ECL
       No Drawings
DRWN
LN.CNT 3729
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to the use of a group of aryl ureas in treating
       raf mediated diseases, and pharmaceutical compositions for use in such
       therapy.
   284461-73-0P 284461-74-1P,
      N-(4-Chloro-3-trifluoromethylphenyl)-N'-[4-[(2-carbamoyl-4-
      pyridyl)oxy]phenyl]urea 284461-82-1P 284461-88-7P
      284462-04-0P 284462-05-1P 284462-17-5P
      284462-18-6P 284462-21-1P 604813-04-9P,
      N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-[3-[5-[[2-
      (dimethylamino)ethyl]carbamoyl]pyridyl]oxy]phenyl]urea
        (preparation of omega-carboxyaryl substituted di-Ph ureas as raf kinase
        inhibitors and anticancer agents)
RN
     284461-73-0 USPATFULL
CN
     2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```

RN 284461-74-1 USPATFULL CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & \\ & & & \\ & & & \\ & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-05-1 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-(CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

RN 604813-04-9 USPATFULL

CN 3-Pyridinecarboxamide, 6-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

```
L15 ANSWER 321 OF 335 USPATFULL on STN
              2003:207917 USPATFULL
ΑN
ΤТ
              Omega-carboxyaryl substituted diphenyl ureas as raf kinase inhibitors
ΙN
              Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF
              Dumas, Jacques, Orange, CT, UNITED STATES
              Khire, Uday, Hamden, CT, UNITED STATES
              Lowinger, Timothy B., Nishinomiya City, JAPAN
              Scott, William J., Guilford, CT, UNITED STATES
              Smith, Roger A., Madison, CT, UNITED STATES
              Wood, Jill E., Hamden, CT, UNITED STATES
              Monahan, Mary-Katherine, Hamden, CT, UNITED STATES
              Natero, Reina, Hamden, CT, UNITED STATES
              Renick, Joel, Milford, CT, UNITED STATES
              Sibley, Robert N., North Haven, CT, UNITED STATES
              BAYER CORPORATION, Pittsburgh, PA, 15205 (non-U.S. corporation)
PΑ
PΙ
              US 20030144278
                                                       A1 20030731
                                                                                                   ABN
ΑI
              US 2002-283248
                                                       A1 20021030 (10)
              Continuation of Ser. No. US 2002-42203, filed on 11 Jan 2002, PENDING
RLI
PRAI
              US 2001-367380P
                                                                20010112 (60)
DT
              Utility
FS
              APPLICATION
              MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
LREP
              1400, ARLINGTON, VA, 22201
              Number of Claims: 67
CLMN
ECL
              Exemplary Claim: 1
DRWN
              No Drawings
LN.CNT 3733
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
              This invention relates to the use of a group of aryl ureas in treating
              raf mediated diseases, and pharmaceutical compositions for use in such
              therapy.
        284461-73-0P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4-[2-(N-1)]-N'-[4
            methylcarbamoyl)-4-pyridyloxy]phenyl]urea 284461-74-1P,
            N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-carbamoyl-4-pyridylox]
            y)phenyl]urea 284461-82-1P 284461-88-7P
            284461-98-9P 284462-04-0P 284462-05-1P
            284462-17-5P 284462-18-6P,
            N-[4-Bromo-3-(trifluoromethyl)phenyl]-N'-[4-[2-(N-methylcarbamoyl)-4-
            pyridyloxy]phenyl]urea 284462-21-1P 474642-55-2P
                 (preparation of diphenylureas as RAF kinase inhibitors)
RN
          284461-73-0 USPATFULL
CN
          2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
               (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
              INDEX NAME)
```

RN 284461-74-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-(CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-05-1 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

RN 474642-55-2 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

```
L15 ANSWER 322 OF 335 USPATFULL on STN
       2003:201617 USPATFULL
AN
       Method and/or process for preparing omega-carboxyaryl substituted
ΤТ
       diphenyl ureas as raf kinas inhibitors
ΙN
       Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF
       Dumas, Jacques, Bethany, CT, UNITED STATES
       Khire, Uday, Hamden, CT, UNITED STATES
       Lowinger, Timothy B., Wuppertal, GERMANY, FEDERAL REPUBLIC OF
       Scott, William J., Guilford, CT, UNITED STATES
       Smith, Roger A., Madison, CT, UNITED STATES
       Wood, Jill E., North Haven, CT, UNITED STATES
PΙ
       US 20030139605
                           A1 20030724
                                              compound claims
       US 7528255
                           B2 20090505
       US 2002-71248
                           A1 20020211 (10)
ΑТ
       Continuation of Ser. No. US 2001-948915, filed on 10 Sep 2001, PENDING
RLI
       Continuation of Ser. No. US 1999-425228, filed on 22 Oct 1999, ABANDONED
       Continuation-in-part of Ser. No. US 1999-257266, filed on 25 Feb 1999,
       ABANDONED
PRAI
       US 1999-115877P
                               19990113 (60)
       US 1999-115878P
                               19990113 (60)
DT
       Utility
FS
       APPLICATION
LREP
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
       Number of Claims: 25
CLMN
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 3287
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to the use of a group of aryl ureas in treating
AΒ
       raf mediated diseases, and pharmaceutical compositions for use in such
       therapy of the formula
       A--D--B wherein
       D is --NH--C(O)--NH--
       A is a substituted moiety of the formula: --L--(M--L.sup.1).sub.q, and
       B is a substituted or unsubstituted up to tricyclic aryl or heteroaryl
       moiety with a t least one 6-member cyclic structure bound directly to D
       containing 0-4 members of the group consisting of nitrogen oxygen and
       sulfur.
       L is a 5-6 membered cyclic structure bound directly to D,
       L.sup.1 comprises a substituted cyclic moiety having at least 5 members
       M is a bridging group having at least one atom and q is an integer of
       from 1-3.
    284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-
      carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P
      284462-17-5P 284462-18-6P
        (preparation of \omega-carboxy(hetero)aryl substituted di-Ph urea raf
        kinase inhibitors by reacting arylisocyanates with arylamines)
RN
     284461-74-1 USPATFULL
     2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
CN
```

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

RN 284462-05-1 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

284462-17-5 USPATFULL RN

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-

(CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl) (CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-(CA INDEX NAME)

L15 ANSWER 323 OF 335 USPATFULL on STN

AN 2003:181526 USPATFULL

TI RAF-MEK-ERK pathway inhibitors to treat cancer

IN Lyons, John F., Moraga, CA, UNITED STATES

Bollag, Gideon, Hercules, CA, UNITED STATES

PI US 20030125359 A1 20030703

US 7307071 B2 20071211

AI US 2002-308721 A1 20021203 (10)

PRAI US 2001-336886P 20011204 (60)

DT Utility

FS APPLICATION

LREP Gregory Giotta, Ph.D., Vice President and Chief Legal Counsel, ONYX Pharmaceuticals, Inc., 3031 Research Drive, Richmond, CA, 94806

CLMN Number of Claims: 11 ECL Exemplary Claim: 1

DRWN 5 Drawing Page(s)

LN.CNT 373

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Materials and methods for treating certain cancers are described, preferably cancers that result from the up-regulation of the RAF-MEK-ERK pathway, and more preferably chronic myelogenous leukemia, and which cancer is preferably resistant to the inhibition of the Bcr-Abl tyrosine kinase, imatinib.

IT 284461-73-0, BAY 43-9006

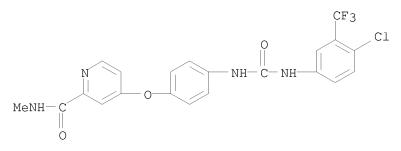
(BAY 43-9006; RAF-MEK-ERK pathway inhibitors to treat cancer)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethy1)pheny1]amino]carbony1]amino]phenoxy]-N-methy1- (CA INDEX NAME)

```
L15 ANSWER 324 OF 335 USPATFULL on STN
       2003:153423 USPATFULL
AN
       Omega-carboxy aryl substituted diphenyl ureas as p38 kinase inhibitors
ΤТ
ΙN
       Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF
       Dumas, Jacques, Orange, CT, UNITED STATES
       Khire, Uday, Handen, CT, UNITED STATES
       Lowinger, Timothy B., Nishinomiya, JAPAN
       William, Scott J., Guilford, CT, UNITED STATES
       Smith, Roger A., Madison, CT, UNITED STATES
       Wood, Jill E., Hamden, CT, UNITED STATES
       Monahan, Mary-Katherine, Hamden, CT, UNITED STATES
       Naero, Reina, Hamden, CT, UNITED STATES
       Renick, Joel, Milford, CT, UNITED STATES
       Sibley, Robert N., North Haven, CT, UNITED STATES
РΤ
       US 20030105091
                           A1 20030605
ΑI
       US 2002-86417
                           A1 20020304 (10)
RLI
       Continuation of Ser. No. US 1999-425229, filed on 22 Oct 1999, ABANDONED
       Continuation-in-part of Ser. No. US 1999-257265, filed on 25 Feb 1999,
       ABANDONED
PRAI
       US 1999-115878P
                               19990113 (60)
DT
       Utility
FS
       APPLICATION
LREP
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
CLMN
       Number of Claims: 38
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 4076
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to the use of a group of aryl ureas in treating
AB
       p38 mediated diseases, and pharmaceutical compositions for use in such
       therapy.
    284461-73-0P 284461-74-1P 284461-82-1P
      284461-88-7P 284461-98-9P 284462-04-0P
      284462-05-1P 284462-17-5P 284462-18-6P
      284462-21-1P
        (preparation of \omega-carboxy aryl substituted di-Ph ureas as p38 kinase
        inhibitors)
RN
     284461-73-0 USPATFULL
CN
     2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```



RN 284461-74-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-05-1 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

$$\begin{array}{c} \text{CF 3} \\ \text{NH-C-NH-C} \\ \text{O} \\ \text{O} \\ \end{array}$$

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

```
L15 ANSWER 325 OF 335 USPATFULL on STN
ΑN
       2002:295343 USPATFULL
       Inhibition of RAF kinase using quinolyl, isoquinolyl or pyridyl ureas
ΤТ
ΙN
      Dumas, Jacques, Orange, CT, UNITED STATES
      Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF
       Khire, Uday, Hamden, CT, UNITED STATES
       Wood, Jill E., Hamden, CT, UNITED STATES
       Robert, Sibley N., North Haven, CT, UNITED STATES
       Monahan, Mary-Katherine, Hamden, CT, UNITED STATES
       Renick, Joel, Milford, CT, UNITED STATES
       Gunn, David E., Hamden, CT, UNITED STATES
       Lowinger, Timothy B., Nishinomiya City, JAPAN
       Scott, William J., Guilford, CT, UNITED STATES
       Smith, Roger A., Madison, CT, UNITED STATES
       BAYER CORPORATION (U.S. corporation)
PA
PΙ
       US 20020165394
                          A1 20021107
                                               no ODP
ΑI
       US 2001-777920
                           A1 20010207 (9)
       Continuation-in-part of Ser. No. US 2001-758548, filed on 12 Jan 2001,
RLI
       PENDING Continuation-in-part of Ser. No. US 1999-425228, filed on 22 Oct
       1999, ABANDONED Continuation-in-part of Ser. No. US 1999-257266, filed
       on 25 Feb 1999, ABANDONED
PRAI
       US 1999-115877P
                               19990113 (60)
DT
       Utility
FS
       APPLICATION
      MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
LREP
       1400, ARLINGTON, VA, 22201
CLMN
      Number of Claims: 33
ECL
      Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 3722
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to the use of a group of aryl ureas in treating
AB
       raf mediated diseases, and pharmaceutical compositions in such therapy.
    284461-73-0P 284461-74-1P 284461-82-1P
      284461-88-7P 284461-98-9P 284462-04-0P
      284462-05-1P 284462-17-5P 284462-18-6P
      284462-21-1P
        (drug candidate; preparation of quinolyl, isoquinolyl or pyridyl-ureas as
        inhibitors of raf kinase)
RN
     284461-73-0 USPATFULL
CN
     2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
       INDEX NAME)
```

RN 284461-74-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & \\ & & & \\ & & & \\ & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl-(CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-05-1 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

IT 474642-55-2

(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

RN 474642-55-2 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

L15 ANSWER 326 OF 335 USPATFULL on STN 2002:262209 USPATFULL ΑN ТΤ Death domain containing receptor-4 antibodies ΙN Ni, Jian, Rockville, MD, United States Rosen, Craig A., Laytonsville, MD, United States Pan, James G., Ypsilanti, MI, United States Gentz, Reiner L., Silver Spring, MD, United States Dixit Wishwa Los Altos Hills, CA, United States (Human Genome Sciences) Inc., Rockville, MD, United States (U.S. PACOLDOLGAtirotal PΙ US 6461823 B1 20021008 ΑТ US 1999-448868 19991124 (9) RLI Division of Ser. No. US 1998-13895, filed on 27 Jan 1998, now patented, Pat. No. US 6342363 US 1997-37829P 19970205 (60) PRAI US 1997-35722P 19970128 (60) DT Utility FS GRANTED EXNAM Primary Examiner: Spector, Lorraine; Assistant Examiner: Kaufman, Claire Sterne, Kessler, Goldstein & Fox, P.L.L.C. LREP CLMN Number of Claims: 146 Exemplary Claim: 135 ECL 13 Drawing Figure(s); 10 Drawing Page(s) LN.CNT 3073 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB The present invention relates to novel Death Domain Containing Receptor-4 (DR4) proteins which are members of the tumor necrosis factor (TNF) receptor family. In particular, isolated nucleic acid molecules are provided encoding the human DR4 proteins. DR4 polypeptides are also provided as are vectors, host cells and recombinant methods for producing the same. The invention further relates to screening methods for identifying agonists and antagonists of DR4 activity. 284461-73-0, BAY 43-9006 (combination chemotherapy with; death domain containing receptor DR4 and methods for inducing apoptosis and treating cancer with DR4 agonist antibodies) RN 284461-73-0 USPATFULL CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA

INDEX NAME)

```
L15 ANSWER 327 OF 335 USPATFULL on STN
ΑN
                2002:251820 USPATFULL
ΤТ
                Carboxyaryl substituted diphenyl ureas as raf kinase inhibitors
ΙN
                Riedl, Bernd, Wuppertal, GERMANY, FEDERAL REPUBLIC OF
               Dumas, Jacques, Ørange, CT, UNITED STATES
               Khire, Uday, Hamden, CT, UNITED STATES
                Lowinger, Timothy B., Nishinomiya City, CANADA
                Scott, William J., Guilford, CT, UNITED STATES
                Smith, Roger A., Madison, CT, UNITED STATES
                Wood, Jill E., Hamden, CT, UNITED STATES
               Monahan, Mary-Katherine, Hamden, CT, UNITED STATES
               Natero, Reina, Hamden, CT, UNITED STATES
                Renick, Joel, San Diego, CA, UNITED STATES
                Sibley, Robert N., North Haven, CT, UNITED STATES
               BAYER CORPORATION, Pittsburgh, PA (non-U.S. corporation)
PA
PТ
               US 20020137774
                                                            A1 20020926
ΑI
               US 2001-907970
                                                            A1
                                                                                                         abn
                                                                      20010719 (9)
               US 1999-115877P
                                                                      19990113 (60)
PRAI
DT
               Utility
FS
               APPLICATION
               MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
LREP
                1400, ARLINGTON, VA, 22201
               Number of Claims: 67
CLMN
ECL
               Exemplary Claim: 1
DRWN
               No Drawings
LN.CNT 3732
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
                This invention relates to the use of a group of aryl ureas in treating
AB
               raf mediated diseases, and pharmaceutical compositions for use in such
               therapy.
         284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[
ΤT
             carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P
             284462-17-5P 284462-18-6P
                  (preparation of \omega-carboxy(hetero)aryl substituted di-Ph urea raf
                  kinase inhibitors by reacting arylisocyanates with arylamines)
RN
           284461-74-1 USPATFULL
CN
           2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
                (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)
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$$\begin{array}{c|c} & & & & \\ & & \\ & & & \\ & & & \\ & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

RN 284462-05-1 USPATFULL
CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl)-(CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl-(CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

L15 ANSWER 328 OF 335 USPATFULL on STN 2002:185638 USPATFULL ΑN ΤТ Death domain containing receptor 5 ΙN Ni, Jian, Rockville, MD, UNITED STATES Gentz, Reiner L., Silver Spring, MD, UNITED STATES Yu, Guo-Liang, Darnestown, MD, UNITED STATES Rossny Craig A. Laytonsville, MD, UNITED STATES Human Genome Sciences Inc. (U.S. corporation) PA PΙ US 20020098550 A1 20020725 US 2001-5842 A1 20011207 (10) ΑI RLI Division of Ser. No. US 1998-42583, filed on 17 Mar 1998, PENDING PRAI US 1997-40846P 19970317 (60) US 1997-54021P 19970729 (60) DT Utility FS APPLICATION LREP STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C., 1100 NEW YORK AVENUE, N.W., SUITE 600, WASHINGTON, DC, 20005-3934 CLMN Number of Claims: 34 ECL Exemplary Claim: 1 DRWN 7 Drawing Page(s) LN.CNT 2876 CAS INDEXING IS AVAILABLE FOR THIS PATENT. The present invention relates to novel Death Domain Containing Receptor-5 (DR5) proteins which are members of the tumor necrosis factor (TNF) receptor family, and have now been shown to bind TRAIL. In particular, isolated nucleic acid molecules are provided encoding the human DR5 proteins. DR5 polypeptides are also provided as are vectors, host cells and recombinant methods for producing the same. The invention further relates to screening methods for identifying antagonists and

IT 284461-73-0, BAY 43-9006

(combination chemotherapy with; DR5-binding agonist antibodies for induction of apoptosis in DR5 expressing cells and for treatment of cancer and hepatitis C virus infections)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

antagonists of DR5 activity.

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

```
L15 ANSWER 329 OF 335 USPATFULL on STN
               2002:78859 USPATFULL
ΑN
               Omega-carboxyaryl substituted diphenyl ureas as raf kinase inhibitors
ΤТ
ΙN
               Uday, Khire, Hamden, CT, UNITED STATES
               Damas, Jacques, Orange, CT, UNITED STATES
             Riedl, Bernd, Muppertal, GERMANY, FEDERAL REPUBLIC OF
               Lowinger, Timothy B., Nishinomiya City, JAPAN
               Scott, William J., Guilford, CT, UNITED STATES
               Smith, Roger A., Madison, CT, UNITED STATES
               Wood, Jill E., Hamden, CT, UNITED STATES
               Monahan, Mary-Katherine, Hamden, CT, UNITED STATES
               Natero, Reina, Hamden, CT, UNITED STATES
               Joel, Renick, Milford, CT, UNITED STATES
               Sibley, Robert N., North Haven, CT, UNITED STATES
               BAYER CORPORATION, Pittsburgh, PA, 15205 (U.S. corporation)
PA
PΙ
               US 20020042517
                                                         A1 20020411
ΑI
               US 2001-948915
                                                         A1
                                                                  20010910 (9)
               Continuation of Ser. No. US 1999-425228, filed on 22 Oct 1999, ABANDONED
RLT
               Continuation-in-part of Ser. No. US 1999-257266, filed on 25 Feb 1999,
               ABANDONED
                                                                                                     abn
PRAI
               US 1999-115877P
                                                                   19990113 (60)
DT
               Utility
FS
               APPLICATION
               MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
LREP
               1400, ARLINGTON, VA, 22201
CLMN
               Number of Claims: 67
ECL
               Exemplary Claim: 1
DRWN
              No Drawings
LN.CNT 3675
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
               This invention relates to the use of a group of aryl ureas in treating
               raf mediated diseases, and pharmaceutical compositions for use in such
        284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[
             carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P
             284462-17-5P 284462-18-6P
                  (preparation of ∞-carboxy(hetero)aryl substituted di-Ph urea raf
                 kinase inhibitors by reacting arylisocyanates with arylamines)
RN
           284461-74-1 USPATFULL
CN
          2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-
               (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)
```

RN 284462-05-1 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-

(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethy1)phenyl]amino]carbonyl]amino]phenoxy]-N-(2-hydroxyethyl) (CA INDEX NAME)

$$\begin{array}{c} \text{CF3} \\ \text{NH-C-NH-C} \\ \text{NH-C-NH-C} \\ \end{array}$$

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CAINDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA
INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

L15 ANSWER 330 OF 335 USPATFULL on STN

AN 2002:19187 USPATFULL

TI Death domain containing receptor 4 nucleic acids and methods

IN Ni, Jian, Rockville, MD, United States

Rosen, Craig A., Laytonsville, MD, United States

Pan, James G., Ypsilanti, MI, United States

Gentz, Reiner L., Silver Spring, MD, United States

Dixit, Vishva M., Los Altos Hills, CA, United States

PA Human Genome Sciences, Inc., Rockville, MD, United States (U.S.

corporation)

PI US 6342363 B1 20020129

AI US 1998-13895 19980127 (9)

PRAI US 1997-35722P 19970128 (60)

US 1997-37829P 19970205 (60)

DT Utility

FS GRANTED

EXNAM Primary Examiner: Spector, Lorraine; Assistant Examiner: Kaufman, Claire M.

LREP Sterne, Kessler, Goldstein & Fox PLLC

CLMN Number of Claims: 303

ECL Exemplary Claim: 286

DRWN 13 Drawing Figure(s); 10 Drawing Page(s)

LN.CNT 3345

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to novel Death Domain Containing Receptor-4 (DR4) proteins which are members of the tumor necrosis factor (TNF) receptor family. In particular, isolated nucleic acid molecules are provided encoding the human DR4 proteins. DR4 polypeptides are also provided as are vectors, host cells and recombinant methods for producing the same. The invention further relates to screening methods for identifying agonists and antagonists of DR4 activity.

IT 284461-73-0, BAY 43-9006

(combination chemotherapy with; death domain containing receptor DR4 and methods for inducing apoptosis and treating cancer with DR4 agonist antibodies)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

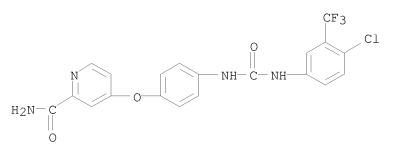
(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CAINDEX NAME)

## COMMAND INTERRUPTED

If this message appears repeatedly, please notify the Help Desk. Enter "HELP STN" for information on contacting the nearest STN Help Desk by telephone or via SEND in the STNMAIL file.

=> d 115 331-335 bib, ab, hitstr

L15 ANSWER 331 OF 335 USPATFULL on STN 2001:188813 USPATFULL AN ΤТ Omega-carboxyaryl substituted diphenyl ureas as raf kinase inhibitors ΙN Riedl, Bernd, Wupperal, Germany, Federal Republic of Dumas, Jacques, Orange, CT, United States Khire, Uday, Hamden, CT, United States Lowinger, Timothy P., Nashnomya City, Japan Scott, William J., Gulford, CT, United States Smith, Roger A., Madison, CT, United States Wood, Jill E., Hamden, CT, United States Monahan, Mary-Katherine, Hamden, CT, United States Natero, Rena, Handen, CT, United States Renick, Joel, Milford, CT, United States Sibley, Robert N., North Haven, CT, United States РΤ US 20010034447 A1 20011025 ΑI US 2001-773604 A1 20010202 (9) RLI Continuation of Ser. No. US 1999-425228, filed on 22 Oct 1999, PENDING Continuation-in-part of Ser. No. US 1999-257266, filed on 25 Feb 1999, ABANDONED PRAI US 1999-115877P 19990113 (60) DT Utility FS APPLICATION LREP MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE 1400, ARLINGTON, VA, 22201 CLMN Number of Claims: 67 ECL Exemplary Claim: 1 DRWN No Drawings LN.CNT 3666 CAS INDEXING IS AVAILABLE FOR THIS PATENT. This invention relates to the use of a group of aryl ureas in treating AB raf mediated diseases, and pharmaceutical compositions for use in such therapy. 284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P 284462-17-5P 284462-18-6P (preparation of ω-carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines) RN 284461-74-1 USPATFULL CN 2-Pyridinecarboxamide, 4-[4-[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)



RN 284462-05-1 USPATFULL
CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethy1)pheny1]amino]carbony1]amino]phenoxy]-N-(2-hydroxyethy1) (CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
INDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

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L15 ANSWER 332 OF 335 USPATFULL on STN
       2001:171152 USPATFULL
AN
       Omega-carboxyaryl substituted disphenyl ureas as raf kinase inhibitors
ΤТ
ΙN
       Riedl, Bernd, Wuppertal, Germany, Federal Republic of
       Dumas, Jaques, Orange, CT, United States
       Khire, Uday, Hamden, CT, United States
       Lowinger, Timothy B., Nishinomiya City, Japan
       Scott, William J., Guilford, CT, United States
       Smith, Roger A., Madison, CT, United States
       Wood, Jill E., Hamden, CT, United States
       Monahan, Mary-Katherine, Hamden, CT, United States
       Natero, Reina, Hamden, CT, United States
       Renick, Joel, Milford, CT, United States
       Sibley, Robert N., Noth Haven, CT, United States
                           A1 20011004
РΤ
       US 20010027202
                                                     abn
ΑI
       US 2001-773658
                           A1 20010202 (9)
RLI
       Continuation of Ser. No. US 1999-425228, filed on 22 Oct 1999, PENDING
       Continuation-in-part of Ser. No. US 1999-257266, filed on 25 Feb 1999,
       ABANDONED
PRAI
       US 1999-115877P
                               19990113 (60)
DT
       Utility
FS
       APPLICATION
LREP
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., Arlington Courthouse Plaza I,
       Suite 1400, 2200 Clarendon Boulevard, Arlington, VA, 22201
       Number of Claims: 67
CLMN
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 3656
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to the use of a group of aryl ureas in treating
AB
       raf mediated diseases, and pharmaceutical compositions for use in such
       therapy.
    284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-
      carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P
      284462-17-5P 284462-18-6P
        (preparation of ω-carboxy(hetero)aryl substituted di-Ph urea raf
        kinase inhibitors by reacting arylisocyanates with arylamines)
RN
     284461-74-1 USPATFULL
     2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
CN
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)
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RN 284462-05-1 USPATFULL
CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethy1)pheny1]amino]carbony1]amino]phenoxy]-N-(2-hydroxyethy1) (CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

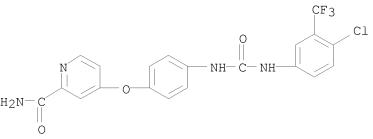
IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

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L15 ANSWER 333 OF 335 USPATFULL on STN
       2001:139616 USPATFULL
AN
ΤТ
       Omega-carboxyaryl substituted diphenyl ureas as raf kinase inhibitors
ΙN
       Riedl, Bernd, Wupperal, Germany, Federal Republic of
       Dumas, Jacques, Orange, CT, United States
       Khire, Uday, Hamden, CT, United States
       Lowinger, Timothy B., Nashnomya City, Japan
       Scott, William J., Gulford, CT, United States
       Smith, Roger A., Madison, CT, United States
       Wood, Jill E., Hamden, CT, United States
       Monahan, Mary-Katherine, Hamden, CT, United States
       Natero, Rena, Hamden, CT, United States
       Renick, Joel, Milford, CT, United States
       Sibley, Robert N., North Haven, CT, United States
                           A1 20010823
РΤ
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ΑI
       US 2001-773672
                           A1 20010202 (9)
RLI
       Continuation of Ser. No. US 1999-425228, filed on 22 Oct 1999, PENDING
       Continuation-in-part of Ser. No. US 1999-257266, filed on 25 Feb 1999,
       ABANDONED
PRAI
       US 1999-115877P
                               19990113 (60)
DT
       Utility
FS
       APPLICATION
LREP
       MILLEN, WHITE, ZELANO & BRANIGAN, P.C., 2200 CLARENDON BLVD., SUITE
       1400, ARLINGTON, VA, 22201
       Number of Claims: 67
CLMN
ECL
       Exemplary Claim: 1
DRWN
      No Drawings
LN.CNT 3652
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention relates to the use of a group of aryl ureas in treating
AR
       raf mediated diseases, and pharmaceutical compositions for use in such
       therapy.
    284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-
      carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P
      284462-17-5P 284462-18-6P
        (preparation of ω-carboxy(hetero)aryl substituted di-Ph urea raf
        kinase inhibitors by reacting arylisocyanates with arylamines)
RN
     284461-74-1 USPATFULL
CN
     2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
       (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)
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RN 284462-05-1 USPATFULL
CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethy1)pheny1]amino]carbony1]amino]phenoxy]-N-(2-hydroxyethy1) (CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
INDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CAINDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

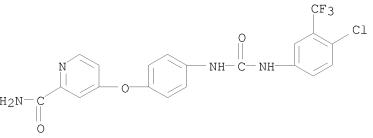
IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

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L15 ANSWER 334 OF 335 USPATFULL on STN
                2001:123628 USPATFULL
AN
ΤТ
                omega-carboxyyaryl substituted diphenyl ureas as raf kinase inhibitors
ΙN
                Riedl, Bernd, Wuppertal, Germany, Federal Republic of
                Dumas, Jacques, Orange, CT, United States
                Khire, Uday, Hamden, CT, United States
                Lowinger, Timothy B., Nishinomiya City, Japan
                Scott, William J., Guilford, CT, United States
                Smith, Roger A., Madison, CT, United States
                Wood, Jill E., Hamden, CT, United States
               Monahan, Mary-Katherine, Hamden, CT, United States
               Natero, Reina, Hamden, CT, United States
                Renick, Joel, Milford, CT, United States
                Sibley, Robert N., North Haven, CT, United States
РΤ
               US 20010011136
                                                            A1 20010802
               US 2001-773675 A1 20010202 (9) abn Continuation of Ser. No. US 1999-425228, filed on 22 Oct 1999, PENDING
ΑI
RLT
                Continuation-in-part of Ser. No. US 1999-257266, filed on 25 Feb 1999,
                ABANDONED
PRAI
               US 1999-115877P
                                                                     19990113 (60)
DT
               Utility
FS
               APPLICATION
LREP
               MILLEN, WHITE, ZELANO & BRANIGAN, P.C., Suite 1400, 2200 Clarendon
               Blvd., Arlington, VA, 22201
               Number of Claims: 67
CLMN
ECL
               Exemplary Claim: 1
DRWN
               No Drawings
LN.CNT 3646
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
               This invention relates to the use of a group of aryl ureas in treating
AB
               raf mediated diseases, and pharmaceutical compositions for use in such
                therapy.
        284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-
             carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P
             284462-17-5P 284462-18-6P
                  (preparation of ω-carboxy(hetero)aryl substituted di-Ph urea raf
                 kinase inhibitors by reacting arylisocyanates with arylamines)
RN
           284461-74-1 USPATFULL
           2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
CN
                (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)
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RN 284462-05-1 USPATFULL
CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethy1)pheny1]amino]carbony1]amino]phenoxy]-N-(2-hydroxyethy1) (CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
INDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CAINDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

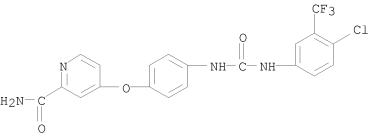
IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

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L15 ANSWER 335 OF 335 USPATFULL on STN
               2001:123627 USPATFULL
AN
ΤТ
               Omega-carboxyaryl subsituted diphenyl ureas as raf kinase inhibitors
ΙN
               Riedl, Bernd, Wuppertal, Germany, Federal Republic of
               Dumas, Jacques, Orange, CT, United States
               Khire, Uday, Hamden, CT, United States
               Lowinger, Timothy B., Nishinomiya City, Japan
               Scott, William J., Guilford, CT, United States
               Smith, Roger A., Madison, CT, United States
               Wood, Jill E., Hamden, CT, United States
               Monahan, Mary-Katherine, Hamden, CT, United States
               Natero, Reina, Hamden, CT, United States
               Renick, Joel, Milford, CT, United States
               Sibley, Robert N., North Haven, CT, United States
РΤ
               US 20010011135
                                                           A1 20010802
               US 2001-773659 A1 20010202 (9) abn
Continuation of Ser. No. US 1999-425228,
ΑI
                                                                                                          filed on 22 Oct 1999, PENDING
RLT
               Continuation-in-part of Ser. No. US 1999-257266, filed on 25 Feb 1999,
               ABANDONED
PRAI
               US 1999-115877P
                                                                     19990113 (60)
DT
               Utility
FS
               APPLICATION
LREP
               MILLEN, WHITE, ZELANO & BRANIGAN, P.C., Suite 1400, Arlington Courthouse
               Plaza 1, Arlington, VA, 22201
               Number of Claims: 67
CLMN
ECL
               Exemplary Claim: 1
DRWN
               No Drawings
LN.CNT 3686
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
               This invention relates to the use of a group of aryl ureas in treating
AB
               raf mediated diseases, and pharmaceutical compositions for use in such
               therapy.
        284461-74-1P, N-[4-Chloro-3-(trifluoromethyl)phenyl]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-284461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-N'-[4-(2-28461-74-1P)]-
             carbamoyl-4-pyridyloxy)phenyl]urea 284462-05-1P
             284462-17-5P 284462-18-6P
                  (preparation of ω-carboxy(hetero)aryl substituted di-Ph urea raf
                 kinase inhibitors by reacting arylisocyanates with arylamines)
RN
           284461-74-1 USPATFULL
           2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-
CN
                (trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]- (CA INDEX NAME)
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RN 284462-05-1 USPATFULL
CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-[2-(dimethylamino)ethyl]- (CA INDEX NAME)

RN 284462-17-5 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethy1)pheny1]amino]carbony1]amino]phenoxy]-N-(2-hydroxyethy1) (CA INDEX NAME)

RN 284462-18-6 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA
INDEX NAME)

IT 284461-82-1P 284461-88-7P 284461-98-9P

284462-04-0P 284462-21-1P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-82-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-

(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CAINDEX NAME)

RN 284461-88-7 USPATFULL

CN 2-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284461-98-9 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)-(CA INDEX NAME)

RN 284462-04-0 USPATFULL

CN 3-Pyridinecarboxamide, 5-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

RN 284462-21-1 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-bromo-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-ethyl- (CA INDEX NAME)

IT 284461-73-0P

(preparation of  $\omega$ -carboxy(hetero)aryl substituted di-Ph urea raf kinase inhibitors by reacting arylisocyanates with arylamines)

RN 284461-73-0 USPATFULL

CN 2-Pyridinecarboxamide, 4-[4-[[[[4-chloro-3-(trifluoromethyl)phenyl]amino]carbonyl]amino]phenoxy]-N-methyl- (CA INDEX NAME)

## 09/993,647

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